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1.2
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YOU HAVE REQUESTED DATA FROM 10 ANSWERS - CONTINUE? Y/(N):y
L2
     ANSWER 1 OF 10 USPATFULL on STN
       2006:158613 USPATFULL
AN
       Poly-gamma-glutamic conjugates for eliciting immune responses directed
TТ
       against bacilli
       Schneerson, Rachel, Bethesda, MD, UNITED STATES
IN
       Leppla, Stephen, Bethesda, MD, UNITED STATES
       Robbins, John B., Chevy Chase, MD, UNITED STATES Shiloach, Joseph, Rockville, MD, UNITED STATES
       Kubler-Kielb, Joanna, Rockville, MD, UNITED STATES
       Liu, Darrell, Bethesda, MD, UNITED STATES
       Majadly, Fathy, Frederick, MD, UNITED STATES
PΙ
       US 2006134143
                           A1
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ΑI
       US 2004-559825
                           A1
                                20040604 (10)
       WO 2004-US17736
                                20040604
                                20051202 PCT 371 date
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       US 2003-476598P
                            20030605 (60)
DT
       Utility
FS
       APPLICATION
       KLARQUIST SPARKMAN, LLP, 121 S.W. SALMON STREET, SUITE #1600, PORTLAND,
LREP
       OR, 97204-2988, US
CLMN
       Number of Claims: 36
       Exemplary Claim: 1
ECL
       2 Drawing Page(s)
DRWN
LN.CNT 2866
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Immunogenic compositions and methods for eliciting an immune response
       against B. anthracis and other bacilli are provided that
       include immunogenic conjugates of a poly-γ-glutamic acid
       (\gamma PGA) polypeptide of B. anthracis, or of another
       Bacillus that expresses a \gammaPGA polypeptide. The \gammaPGA
       conjugates elicit an effective immune response against B.
       anthracis, or against another Bacillus, in mammalian hosts to
       which the conjugates are administered.
     ANSWER 2 OF 10 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
L2
     DUPLICATE 1
     2006:436213 BIOSIS
ΔN
     PREV200600430224
DN
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- TI Additional conjugation methods and immunogenicity of Bacillus anthracis poly-gamma-D-glutarnic acid-protein conjugates.
- AU Kubler-Kielb, Joanna [Reprint Author]; Liu, Teh-Yung; Mocca, Christopher; Majadly, Fathy; Robbins, John B.; Schneerson, Rachel
- CS NICHD, NIH, LDMI, 9000 Rockville Pike, Bldg 6, Rm 1A05, Bethesda, MD 20892 USA
- kielbj@mail.nih.gov
- SO Infection and Immunity, (AUG 2006) Vol. 74, No. 8, pp. 4744-4749. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 30 Aug 2006 Last Updated on STN: 30 Aug 2006
- AB The capsule of Bacillus anthracis, composed of poly-gamma-D-glutamic acid (gamma DPGA), is an essential virulence factor of B. anthracis. The capsule inhibits innate host defense through its antiphagocytic action. gamma DPGA is a poor immunogen, but when covalently bound to a carrier protein, it elicits serum antibodies. To identify the optimal construct for clinical use, synthetic gamma DPGAs of different lengths were bound to carrier proteins at different densities. The advantages of the synthetic over the natural polypeptide are the homogeneous chain length and end groups, allowing conjugates to be accurately characterized and standardized and their chemical compositions to be related to their immunogenicities. In the present study, we evaluated, in addition to methods reported by us, hydrazone, oxime, and thioether linkages between gamma DPGA and several proteins, including bovine serum albumin, recombinant Pseudomonas aeruginosa exotoxin A, recombinant B. anthracis protective antigen (rPA), and tetanus toxoid (TT). The effects of the dosage and formulation on the immunogenicities of the conjugates were evaluated in mice. All conjugates were immunogenic. The optimal gamma DPGA chain length of 10 to 15 amino acids and the density, an average of 15 mol gamma DPGA per mol of protein, were confirmed. The thioether bond was the optimal linkage type, and TT and rPA were the best carriers. The optimal dosage was 1.2 to 2.5 mu g of gamma DPGA per mouse, and adsorption of the conjugates onto aluminum hydroxide significantly increased the antibody response to the protein with a lesser effect on anti-gamma DPGA levels.
- L2 ANSWER 3 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2
- AN 2005:1294042 CAPLUS
- DN 144:35295
- TI Hydrazone conjugates of haptens and antigens
- IN Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy; Leppla, Stephen H.; Robbins, John B.; Liu, Darrell T.; Shiloah, Joseph
- PA United States Dept. of Health and Human Services, USA
- SO U.S. Pat. Appl. Publ., 17 pp., Cont.-in-part of Appl. No. PCT/US04/017736. CODEN: USXXCO
- DT Patent
- LA English
- FAN.CNT 3

	PATENT NO.					KIND DATE			7	APPLICATION NO.					DATE				
PI	US 2005271675 WO 2005000884 WO 2005000884				A1 20051208 A1 20050106 C1 20051006				US 2			20041206 20040604							
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                                 20040604
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                           P
                                 20030605
     US 2004-5851
                           Α
                                 20041206
     The authors disclose methods for making an immunogenic conjugate that
AB
     includes a hapten or an antigen covalently linked to a carrier.
     methods include reacting a first agent with a dihydrazide resulting in a
     hydrazino-modified first agent, wherein the first agent is a hapten, an
     antigen or a carrier; reacting a second agent with a benzaldehyde compound
     resulting in a benzaldehyde-modified second agent, wherein the second
     agent is a hapten, an antigen or a carrier, provided that the first agent
     or the second agent is a carrier; and reacting the hydrazine-modified
     first agent with the benzaldehyde-modified second agent resulting in an
     immunogenic conjugate comprising a hapten or an antigen covalently linked
     to a carrier via a hydrazone linkage.
L2
     ANSWER 4 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN
AN
     2005:1314018 CAPLUS
     144:35300
DN
     Methods for preparing immunogenic conjugates for use in vaccines
TI
     Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy;
IN
     Leppla, Stephen H.; Robbins, John B.; Liu, Darrell T.; Shiloach, Joseph
PA
     The Government of the United States of America as Represented by the
     Secretary, Department of Healthand Human Services, USA
     PCT Int. Appl., 53 pp.
SO
     CODEN: PIXXD2
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     English
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    US 2004-5851
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    US 2003-476598P
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                              20030605
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os
    Methods for making an immunogenic conjugate that includes a hapten or an
AB
    antigen covalently linked to a carrier are discussed. The methods include
    reacting a first agent with a dihydrazide resulting in a
    hydrazino-modified first agent, wherein the first agent is a hapten, an
    antigen or a carrier; reacting a second agent with a benzaldehyde compound
    resulting in a benzaldehyde-modified second agent, wherein the second
    agent is a hapten, an antigen or a carrier, provided that the first agent
    or the second agent is a carrier; and reacting the hydrazine-modified
    first agent with the benzaldehyde-modified second agent resulting in an
    immunogenic conjugate comprising a hapten or an antigen covalently linked
    to a carrier via a hydrazone linkage. The examples discuss the
    conjugation of Bacillus poly-\gamma-glutamic acids to carriers such as
    bovine serum albumin, Bacillus anthracis protective antigen, and
    Pseudomonas aeruginosa exotoxin A. The conjugates were shown to induce
    IgG and opsonophagocytosis.
             THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 4
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
L2
    ANSWER 5 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN
AN
    2005:14426 CAPLUS
DN
    142:112426
ΤI
    Bacillus capsular poly-γ-glutamic acid conjugates for eliciting
    immune responses against Bacillus infection
    Schneerson, Rachel; Leppla, Stephen; Robbins, John B.; Shiloach,
IN
    Joseph; Kubler-Kielb, Joanna; Liu, Darrell; Majadly, Fathy
PA
    United States Dept. of Health and Human Services, USA
SO
    PCT Int. Appl., 67 pp.
    CODEN: PIXXD2
DT
    Patent
LA
    English
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PRAI US 2003-476598P
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     WO 2004-US17736
                        W
                                20040604
     US 2004-5851
                         Α
                                20041206
AB
     Immunogenic compns. and methods for eliciting an immune response against
     B. anthracis and other bacilli are provided that include
     immunogenic conjugates of a poly-\gamma-glutamic acid (\gammaPGA)
     polypeptides of B. anthracis, or of another Bacillus that
     expresses a \gammaPGA polypeptide. The \gammaPGA conjugates elicit an
     effective immune response against B. anthracis, or against
     another Bacillus, in mammalian hosts to which the conjugates are
     administered. The conjugate consists of \gamma-D-PGA and carrier
     selected from bovine serum albumin, recombinant Bacillus protective
     antigen, recombinant Pseudomonas aeruginosa exotoxin A, tetanus toxoid,
     diphtheria toxoid, pertussis toxoid, Clostridium perfringens toxoid,
     HBsAg, HBcAg, heyhole limpet hemocyanin, horseshoe crab hemocyanin,
     edestin, mammalian serum albumin, mammalian Ig., analog or mimetic, or
     combination of two or more. The preferred conjugate consists of
     \gamma\text{-D-PGA} and Bacillus protective antigen.
              THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 9
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 6 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 3
L2
     2004:331575 CAPLUS
AN
     140:338027
DN
ΤI
     Methods for preparing Bacillus anthracis protective antigen for
     use in vaccines
     Shiloach, Joseph; Leppla, Stephen H.; Ramirez, Delia M.; Schneerson,
IN
     Rachel; Robbins, John B.
PA
so
     U.S. Pat. Appl. Publ., 13 pp.
     CODEN: USXXCO
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     Patent
LA
     English
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     US 2004076638
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                                                                   20021108
PRAI US 2001-344505P
                         P
                                20011109
     The authors disclose improved methods of producing and recovering B.
     anthracis protective antigen (PA), especially modified PA which is
     protease resistant, and to methods of using of these PAs or nucleic acids
     encoding these PAs for eliciting an immunogenic response.
     ANSWER 7 OF 10 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
L2
     2005:488548 BIOSIS
AN
DN
     PREV200510291216
ΤI
     Future vaccine development at NICHD.
AU
    Robbins, John B. [Reprint Author]; Schneerson, Rachel
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NICHD, Lab Dev and Mol Immun, Sect Bacterial Dis Pathogenesis and Immun,

NIH, Bldg 6, Room 436, Bethesda, MD 20892 USA

CS

robbinsj@nichd.nih.gov; schneerr@mail.nih.gov

SO Kaler, SG [Editor]; Rennert, OM [Editor]. (2004) pp. 49-59. Annals of the New York Academy of Sciences.

Publisher: NEW YORK ACAD SCIENCES, 2 EAST 63RD ST, NEW YORK, NY 10021 USA. Series: ANNALS OF THE NEW YORK ACADEMY OF SCIENCES.

Meeting Info.: 40th Scientific Symposium of the National-Institute-of-Child-Health-and-Human-Development. Bethesda, MD, USA. 20030908,. NICHD. ISSN: 0077-8923 (print). ISBN: 1-57331-520-6(H).

DT Book; (Book Chapter)
Conference; (Meeting)
General Review; (Literature Review)

LA English

ED Entered STN: 16 Nov 2005 Last Updated on STN: 16 Nov 2005

- Using published data and the results of our studies, we hypothesized that a critical level of serum IgG antibodies to the surface structures of invasive pathogens (capsular polysaccharides of Haemophilus influenzae type b, pneumococcus, meningococcus, Salmonella typhi, Escherichia coli, and Staphylococcus aureus, the O-specific polysaccharide LPS domain of the LPS of Shigella, non-typhoidal Salmonella, and E. coli, and the capsular polypeptide of Bacillus anthraces) confer immunity to these pathogens. Covalent attachment to a protein increases their immunogenicity and bestows T-cell properties to these antigens. We have also shown that a critical level of serum IgG antibodies to pertussis toxin alone induces immunity on both an individual and on a community basis (herd immunity) to Bordetella pertussis. It is likely that all the above conjugates and pertussis toxoid will be incorporated into vaccines for routine infant immunization.
- L2 ANSWER 8 OF 10 MEDLINE on STN
- AN 2005202078 MEDLINE
- DN PubMed ID: 15838097
- TI Future vaccine development at NICHD.
- AU Robbins John B; Schneerson Rachel
- CS Laboratory of Developmental and Molecular Immunity, NICHD, NIH, Building 6, Room 436, Bethesda, MD 20892, USA.. robbinsj@nichd.nih.gov
- SO Annals of the New York Academy of Sciences, (2004 Dec) Vol. 1038, pp. 49-59.

Journal code: 7506858. ISSN: 0077-8923.

- CY United States
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Priority Journals
- EM 200511
- ED Entered STN: 20 Apr 2005 Last Updated on STN: 10 Nov 2005 Entered Medline: 9 Nov 2005
- AB Using published data and the results of our studies, we hypothesized that a critical level of serum IgG antibodies to the surface structures of invasive pathogens (capsular polysaccharides of Haemophilus influenzae type b, pneumococcus, meningococcus, Salmonella typhi, Escherichia coli, and Staphylococcus aureus, the O-specific polysaccharide LPS domain of the LPS of Shigella, non-typhoidal Salmonella, and E. coli, and the capsular polypeptide of Bacillus anthraces) confer immunity to these pathogens. Covalent attachment to a protein increases their immunogenicity and bestows T-cell properties to these antigens. We have also shown that a critical level of serum IgG antibodies to pertussis toxin alone induces immunity on both an individual and on a community basis (herd immunity) to Bordetella pertussis. It is likely that all the above conjugates and pertussis toxoid will be incorporated into vaccines for routine infant immunization.
- L2 ANSWER 9 OF 10 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN DUPLICATE 4

- AN 2003:478120 BIOSIS
- DN PREV200300478120
- TI Poly(gamma-D-glutamic acid) protein conjugates induce IgG antibodies in mice to the capsule of Bacillus anthracis: A potential addition to the anthrax vaccine.
- AU Schneerson, Rachel [Reprint Author]; Kubler-Kielb, Joanna; Liu, Teh-Yung; Dai, Zhong-Dong; Leppla, Stephen H.; Yergey, Alfred; Backlund, Peter; Shiloach, Joseph; Majadly, Fathy; Robbins, John B.
- CS National Institute of Child Health and Human Development, National Institutes of Health, Bethesda, MD, 20892, USA schneerr@mail.nih.gov
- SO Proceedings of the National Academy of Sciences of the United States of America, (July 22 2003) Vol. 100, No. 15, pp. 8945-8950. print. ISSN: 0027-8424 (ISSN print).
- DT Article
- LA English
- ED Entered STN: 15 Oct 2003 Last Updated on STN: 15 Oct 2003
- AB Both the protective antigen (PA) and the poly(gamma-D-glutamic acid) capsule (gammaDPGA) are essential for the virulence of Bacillus anthracis. A critical level of vaccine-induced IgG anti-PA confers immunity to anthrax, but there is no information about the protective action of IgG anti-gammaDPGA. Because the number of spores presented by bioterrorists might be greater than encountered in nature, we sought to induce capsular antibodies to expand the immunity conferred by available anthrax vaccines. The nonimmunogenic gammaDPGA or corresponding synthetic peptides were bound to BSA, recombinant B. anthracis PA (rPA), or recombinant Pseudomonas aeruginosa exotoxin A (rEPA). To identify the optimal construct, conjugates of B. anthracis gammaDPGA, Bacillus pumilus gammaDLPGA, and peptides of varying lengths (5-, 10-, or 20-mers), of the D or L configuration with active groups at the N or C termini, were bound at 5-32 mol per protein. The conjugates were characterized by physico-chemical and immunological assays, including GLC-MS and matrix-assisted laser desorption ionization time-of-flight spectrometry, and immunogenicity in 5- to 6-week-old mice. IgG anti-gammaDPGA and antiprotein were measured by ELISA. The highest levels of IgG anti-gammaDPGA were elicited by decamers of gammaDPGA at 10-20 mol per protein bound to the N- or C-terminal end. High IgG anti-gammaDPGA levels were elicited by two injections of 2.5 mug of gammaDPGA per mouse, whereas three injections were needed to achieve high levels of protein antibodies. rPA was the most effective carrier. Anti-gammaDPGA induced opsonophagocytic killing of B. anthracis tox-, cap+. gammaDPGA conjugates may enhance the protection conferred by PA alone. gammaDPGA-rPA conjugates induced both anti-PA and anti-gammaDPGA.
- L2 ANSWER 10 OF 10 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN DUPLICATE 5
- AN 2002:429815 BIOSIS
- DN PREV200200429815
- TI Development of an improved vaccine for anthrax.
- AU Leppla, Stephen H. [Reprint author]; Robbins, John B.; Schneerson, Rachel; Shiloach, Joseph
- CS Oral Infection and Immunity Branch, NIDCR, 30 Convent Drive, Building 30, Room 303, Bethesda, MD, 20892-4350, USA Leppla@nih.gov
- SO Journal of Clinical Investigation, (July, 2002) Vol. 110, No. 2, pp. 141-144. print.

 CODEN: JCINAO. ISSN: 0021-9738.
- DT Article
- LA English
- ED Entered STN: 14 Aug 2002 Last Updated on STN: 14 Aug 2002

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E5
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E6
            1
                   LEPPLA STEVE H/AU
E7
            36
                   LEPPLA W/AU
            9
                   LEPPLA WOLFRAM/AU
E8
E9
            3
                   LEPPLA WOLLSIFFER G/AU
E10
            1
                   LEPPLAWOLLSIFFER G/AU
E11
             3
                   LEPPLE A P/AU
E12
             1
                   LEPPLE ALBRECHT P/AU
=> s e1-e6 and anthra?
           309 ("LEPPLA S M"/AU OR "LEPPLA STEPHAN H"/AU OR "LEPPLA STEPHEN"/AU
                OR "LEPPLA STEPHEN A"/AU OR "LEPPLA STEPHEN H"/AU OR "LEPPLA
              STEVE H"/AU) AND ANTHRA?
=> s 13 and glutamic
            18 L3 AND GLUTAMIC
=> dup rem 14
PROCESSING COMPLETED FOR L4
             15 DUP REM L4 (3 DUPLICATES REMOVED)
=> d bib ab 1-
YOU HAVE REQUESTED DATA FROM 15 ANSWERS - CONTINUE? Y/(N):y
1.5
     ANSWER 1 OF 15 USPATFULL on STN
AN
       2006:158613 USPATFULL
TΙ
       Poly-gamma-glutamic conjugates for eliciting immune responses
       directed against bacilli
IN
       Schneerson, Rachel, Bethesda, MD, UNITED STATES
         Leppla, Stephen, Bethesda, MD, UNITED STATES
       Robbins, John B., Chevy Chase, MD, UNITED STATES
       Shiloach, Joseph, Rockville, MD, UNITED STATES
       Kubler-Kielb, Joanna, Rockville, MD, UNITED STATES
       Liu, Darrell, Bethesda, MD, UNITED STATES
       Majadly, Fathy, Frederick, MD, UNITED STATES
PΙ
       US 2006134143
                               20060622
                          Α1
ΑI
       US 2004-559825
                          Α1
                               20040604 (10)
       WO 2004-US17736
                               20040604
                               20051202 PCT 371 date
PRAI
       US 2003-476598P
                           20030605 (60)
DT
       Utility
FS
       APPLICATION
       KLARQUIST SPARKMAN, LLP, 121 S.W. SALMON STREET, SUITE #1600, PORTLAND,
LREP
       OR, 97204-2988, US
CLMN
       Number of Claims: 36
ECL
       Exemplary Claim: 1
DRWN
       2 Drawing Page(s)
LN.CNT 2866
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB
       Immunogenic compositions and methods for eliciting an immune response
       against B. anthracis and other bacilli are provided that
       include immunogenic conjugates of a poly-γ- glutamic acid
       (γPGA) polypeptide of B. anthracis, or of another
       Bacillus that expresses a \gammaPGA polypeptide. The \gammaPGA
       conjugates elicit an effective immune response against B.
       anthracis, or against another Bacillus, in mammalian hosts to
       which the conjugates are administered.
```

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L5
     ANSWER 2 OF 15 USPATFULL on STN
       2006:41161 USPATFULL
AN
       Methods and formulations comprising agonists and antagonists of nuclear
TI
       hormone receptors
       Sternberg, Esther M., 3610 UPTON AVENUE N.W., WASHINGTON, DC, UNITED
IN
       STATES 20008
       Webster, Jeannette I., Washington, DC, UNITED STATES
       Tonelli, Leonardo H., Bethesda, MD, UNITED STATES
        Leppla, Stephen H., Bethesda, MD, UNITED STATES
       Moayeri, Mahtab, Bethesda, MD, UNITED STATES
       US 2006035813
                          A1
                               20060216
ΡI
       US 2003-530254
                               20031003 (10)
ΑI
                          A1
       WO 2003-US31406
                               20031003
                               20050404 PCT 371 date
PRAI
       US 2002-416222P
                           20021004 (60)
       US 2003-419454P
                           20021018 (60)
DT
       Utility
FS
       APPLICATION
       KLARQUIST SPARKMAN, LLP, 121 S.W. SALMON STREET, SUITE #1600, ONE WORLD
LREP
       TRADE CENTER, PORTLAND, OR, 97204-2988, US
CLMN
      Number of Claims: 51
       Exemplary Claim: 1
ECL
       12 Drawing Page(s)
DRWN
LN.CNT 4767
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Novel compounds, pharmaceutical compositions, and methods are provided
       for modulating processes mediated by nuclear hormone receptors. A
       partial or complete agonist or antagonist modulates, directly or
       indirectly, an activity of one or more nuclear hormone receptors for
       glucocorticoids (GRs), androgens (ARs), mineralocorticoids (MRs),
      progestins (PRs), estrogens (ERs), thyroid hormones (TRs), vitamin D
       (VDRs), retinoids (RARs and RXRs), peroxisomes (XPARs and PPARs),
       icosanoids (IRs), or one or more orphan receptors, such as steroid and
       thyroid receptors. Exemplary compounds of the disclosure are bacterial
       products, for example bacterial toxins, and these compounds are useful
       in screens for other antagonists and agonists. Related methods and
       compositions are provided for diagnosis, treatment and prevention of
       bacterial disease and associated or unrelated inflammatory, autoimmune,
       toxic (including shock), and chronic and/or lethal sequelae associated
       with bacterial infection.
     ANSWER 3 OF 15 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 1
L5
     2005:1294042 CAPLUS
AN
DN
     144:35295
     Hydrazone conjugates of haptens and antigens
ΤI
     Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy; Leppla,
IN
     Stephen H.; Robbins, John B.; Liu, Darrell T.; Shiloah, Joseph
PA
     United States Dept. of Health and Human Services, USA
SO
     U.S. Pat. Appl. Publ., 17 pp., Cont.-in-part of Appl. No. PCT/US04/017736.
     CODEN: USXXCO
DT
     Patent
LA
     English
FAN.CNT 3
                                           APPLICATION NO.
     PATENT NO.
                        KIND
                               DATE
                                                                   DATE
                        ----
                                -----
                                           -----
PΙ
    US 2005271675
                         A1
                               20051208
                                           US 2004-5851
                                                                   20041206
                         A1
     WO 2005000884
                               20050106
                                           WO 2004-US17736
                                                                   20040604
                               20051006
     WO 2005000884
                         C1
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            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
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TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

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             EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
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             SN, TD, TG
     WO 2005117965
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                                20051215
                                            WO 2005-US19678
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             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
             LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
             NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK,
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             RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
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PRAI WO 2004-US17736
                         A2
                                20040604
     US 2003-476598P
                          Ρ
                                20030605
     US 2004-5851
                          A
                                20041206
     The authors disclose methods for making an immunogenic conjugate that
AB
     includes a hapten or an antigen covalently linked to a carrier.
     methods include reacting a first agent with a dihydrazide resulting in a
     hydrazino-modified first agent, wherein the first agent is a hapten, an
     antiqen or a carrier; reacting a second agent with a benzaldehyde compound
     resulting in a benzaldehyde-modified second agent, wherein the second
     agent is a hapten, an antigen or a carrier, provided that the first agent
     or the second agent is a carrier; and reacting the hydrazine-modified
     first agent with the benzaldehyde-modified second agent resulting in an
     immunogenic conjugate comprising a hapten or an antigen covalently linked
     to a carrier via a hydrazone linkage.
     ANSWER 4 OF 15 CAPLUS COPYRIGHT 2006 ACS on STN
L5
AN
     2005:1314018 CAPLUS
     144:35300
DN
     Methods for preparing immunogenic conjugates for use in vaccines
TI
     Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy; Leppla,
IN
     Stephen H.; Robbins, John B.; Liu, Darrell T.; Shiloach, Joseph
     The Government of the United States of America as Represented by the
PA
     Secretary, Department of Healthand Human Services, USA
so
     PCT Int. Appl., 53 pp.
     CODEN: PIXXD2
     Patent
DT
LA
    English
FAN.CNT 3
     PATENT NO.
                         KIND
                                DATE
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                                                                   DATE
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                                                                   20050603
PΙ
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             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT,
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             MR, NE, SN, TD, TG
     WO 2005000884
                          A1
                                20050106
                                            WO 2004-US17736
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                          C1
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        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
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CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
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             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
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             EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
             SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
             SN, TD, TG
     US 2005271675
                          A1
                                 20051208
                                             US 2004-5851
                                                                     20041206
PRAI WO 2004-US17736
                          A
                                 20040604
     US 2004-5851
                          Α
                                 20041206
     US 2003-476598P
                          Ρ
                                 20030605
     MARPAT 144:35300
OS
     Methods for making an immunogenic conjugate that includes a hapten or an
AB
     antigen covalently linked to a carrier are discussed.
                                                             The methods include
     reacting a first agent with a dihydrazide resulting in a
     hydrazino-modified first agent, wherein the first agent is a hapten, an
     antigen or a carrier; reacting a second agent with a benzaldehyde compound
     resulting in a benzaldehyde-modified second agent, wherein the second
     agent is a hapten, an antigen or a carrier, provided that the first agent
     or the second agent is a carrier; and reacting the hydrazine-modified
     first agent with the benzaldehyde-modified second agent resulting in an
     immunogenic conjugate comprising a hapten or an antigen covalently linked
     to a carrier via a hydrazone linkage. The examples discuss the
     conjugation of Bacillus poly-\gamma- glutamic acids to carriers
     such as bovine serum albumin, Bacillus anthracis protective
     antigen, and Pseudomonas aeruginosa exotoxin A. The conjugates were shown
     to induce IgG and opsonophagocytosis.
              THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 4
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L5
     ANSWER 5 OF 15 CAPLUS COPYRIGHT 2006 ACS on STN
AN
     2005:14426 CAPLUS
     142:112426
DN
     Bacillus capsular poly-γ- glutamic acid conjugates for
TI
     eliciting immune responses against Bacillus infection
     Schneerson, Rachel; Leppla, Stephen; Robbins, John B.; Shiloach,
IN
     Joseph; Kubler-Kielb, Joanna; Liu, Darrell; Majadly, Fathy
     United States Dept. of Health and Human Services, USA
PA
SO
     PCT Int. Appl., 67 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 3
     PATENT NO.
                         KIND
                                DATE
                                             APPLICATION NO.
                                                                    DATE
                         _ _ _ _
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PΙ
     WO 2005000884
                          A1
                                20050106
                                             WO 2004-US17736
                                                                    20040604
                          C1
                                20051006
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             AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
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             SN, TD, TG
     AU 2004252091
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                                             AU 2004-252091
                                                                     20040604
     CA 2528067
                          AΑ
                                20050106
                                             CA 2004-2528067
                                                                    20040604
     EP 1633778
                          A1
                                20060315
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                                                                     20040604
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R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
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                                20051208
                                            US 2004-5851
                                                                     20041206
     US 2005271675
                          A1
                                20051215
                                             WO 2005-US19678
     WO 2005117965
                          Α1
                                                                     20050603
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             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
             LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
             NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK,
             SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU,
             ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
             EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT,
             RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
             MR, NE, SN, TD, TG
     US 2006134143
                          A1
                                20060622
                                            US 2005-559825
                                                                     20051202
PRAI US 2003-476598P
                          Ρ
                                20030605
                          W
     WO 2004-US17736
                                20040604
     US 2004-5851
                          Α
                                20041206
AB
     Immunogenic compns. and methods for eliciting an immune response against
     B. anthracis and other bacilli are provided that include
     immunogenic conjugates of a poly-γ- glutamic acid
     (\gamma PGA) polypeptides of B. anthracis, or of another
     Bacillus that expresses a \gammaPGA polypeptide. The \gammaPGA
     conjugates elicit an effective immune response against B.
     anthracis, or against another Bacillus, in mammalian hosts to
     which the conjugates are administered. The conjugate consists of
     \gamma-D-PGA and carrier selected from bovine serum albumin, recombinant
     Bacillus protective antigen, recombinant Pseudomonas aeruginosa exotoxin
     A, tetanus toxoid, diphtheria toxoid, pertussis toxoid, Clostridium
     perfringens toxoid, HBsAg, HBcAg, heyhole limpet hemocyanin, horseshoe
     crab hemocyanin, edestin, mammalian serum albumin, mammalian Ig., analog
     or mimetic, or combination of two or more. The preferred conjugate
     consists of \gamma-D-PGA and Bacillus protective antigen.
              THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 9
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 6 OF 15 USPATFULL on STN
L5
       2005:292537 USPATFULL
ΑN
       Multimeric protein toxins to target cells having multiple identifying
TΤ
       characteristics
IN
       Leppla, Stephen H., Bethesda, MD, UNITED STATES
       Liu, Shi-Hui, Gaithersburg, MD, UNITED STATES
       Bugge, Thomas H., Bethesda, MD, UNITED STATES
       The Government of the United States, as represented by the secretary of
PA
       health and Human (U.S. corporation)
       Services, National Institutes of Health, Office of Technology Transfer,
       Rockville, MD, UNITED STATES (U.S. corporation)
PΙ
       US 2005255083
                          A1
                               20051117
       US 2005-55557
ΑI
                          A1
                               20050209 (11)
       US 2004-543417P
PRAI
                           20040209 (60)
DT
       Utility
       APPLICATION
FS
       TOWNSEND AND TOWNSEND AND CREW, LLP, TWO EMBARCADERO CENTER, 8TH FLOOR,
LREP
       SAN FRANCISCO, CA, 94111, US
CLMN
       Number of Claims: 46
ECL
       Exemplary Claim: 1
DRWN
       8 Drawing Page(s)
LN.CNT 4299
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention provides compositions comprising modified
       bacterial toxins and methods for using the modified bacterial toxins for
```

targeting particular cell populations and for treating diseases.

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L5
     ANSWER 7 OF 15 USPATFULL on STN
AN
       2005:226972 USPATFULL
TI
       Anthrax lethal factor is a mapk kinase protease
       Duesbery, Nicholas, Grand Rapids, MI, UNITED STATES
IN
       Webb, Craig, Rockford, MI, UNITED STATES
         Leppla, Stephen, Bethesda, MD, UNITED STATES
       Vande Woude, George, Ada, MI, UNITED STATES
PA
       The Gov. of the USA as represented by the Secretary of the Dept of
       Health and Human Services, Rockville, MD, UNITED STATES (U.S.
       corporation)
       US 2005196822
                          A1
                               20050908
PΙ
       US 7056693
                               20060606
                          B2
AΤ
       US 2005-112137
                          A1
                               20050422 (11)
       Division of Ser. No. US 2002-93200, filed on 5 Mar 2002, GRANTED, Pat.
RLI
       No. US 6911203 Division of Ser. No. US 2000-623104, filed on 13 Dec
       2000, GRANTED, Pat. No. US 6485925 A 371 of International Ser. No. WO
       1999-US7129, filed on 31 Mar 1999
PRAI
       US 1998-80330P
                           19980401 (60)
DT
       Utility
FS
       APPLICATION
       TOWNSEND AND TOWNSEND AND CREW, LLP, TWO EMBARCADERO CENTER, 8TH FLOOR,
LREP
       SAN FRANCISCO, CA, 94111, US
CLMN
       Number of Claims: 8
       Exemplary Claim: 1-24
ECL
DRWN
       1 Drawing Page(s)
LN.CNT 2431
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention relates to in vitro and ex vivo methods of
AB
       screening for modulators, homologues, and mimetics of lethal factor
       mitogen activated protein kinase kinase (MAPKK) protease activity, as
       well as methods of treating cancer by administering LF to transformed
       cells.
L5
     ANSWER 8 OF 15 USPATFULL on STN
       2005:143741 USPATFULL
AN
       Imaging the activity of extracellular protease in cells using mutant
TΙ
       anthrax toxin protective antigens that are cleaved by specific
       extracellular proteases
       Bugge, Thomas H., Bethesda, MD, UNITED STATES
IN
         Leppla, Stephen H., Bethesda, MD, UNITED STATES
       Liu, Shi-Hui, Rockville, MD, UNITED STATES
       Mitola, David, Baltimore, MD, UNITED STATES
PA
       The Government of the United States as represented by the Secretary of
       the Department of Health and, Rockville, MD, UNITED STATES, 20852-3804
       (U.S. corporation)
PΙ
       US 2005123476
                          A1
                               20050609
       US 2003-488806
ΑI
                          A1
                               20020905 (10)
       WO 2002-US28397
                               20020905
PRAI
       US 2001-317550P
                           20010905 (60)
DT
       Utility
FS
       APPLICATION
LREP
       TOWNSEND AND TOWNSEND AND CREW, LLP, TWO EMBARCADERO CENTER, 8TH FLOOR,
       SAN FRANCISCO, CA, 94111, US
CLMN
       Number of Claims: 28
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 4268
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       This invention pertains to methods for imaging the activity of
       extracellular proteases in cells using the anthrax binary
       toxin-system to target cells expressing extracellular proteases with
       mutant anthrax toxin protective antigens (µPrAg) that bind
       to receptors on the cells and are cleaved by a specific extracellular
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protease expressed by the cells, and ligands that specifically bind to the cleaved $\mu PrAg$ and are linked to a moiety that is detectable by an imaging procedure. The $\mu PrAg$ proteins used in the methods comprise a protease cleavage site that is cleaved by a specific extracellular protease and is in place of the furin cleavage site of the native PrAg. The methods are useful for diagnosing and treating diseases and undesirable physiological conditions correlated with the activity of extracellular proteases, and for optimizing the therapeutic efficacy of drugs used to treat such diseases and conditions.

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ANSWER 9 OF 15 USPATFULL on STN
L5
      2004:221352 USPATFULL
AN
      Methods for preparing Bacillus anthracis sporulation deficient
ΤI
      mutants and for producing recombinant Bacillus anthracis
      protective antigen for use in vaccines
      Leppla, Stephen H., Bethesda, MD, UNITED STATES
TN
      Rosovitz, Mary Jo, Kensington, MD, UNITED STATES
      Hsu, S. Dana, Bethesda, MD, UNITED STATES
PΤ
      US 2004171121
                         A1
                               20040902
ΑI
      US 2003-638006
                         A1
                               20030808 (10)
PRAI
      US 2002-402285P
                          20020809 (60)
DT
      Utility
      APPLICATION
FS
      KLARQUIST SPARKMAN, LLP, 121 S.W. SALMON STREET, SUITE #1600, ONE WORLD
LREP
      TRADE CENTER, PORTLAND, OR, 97204-2988
      Number of Claims: 67
CLMN
ECL
      Exemplary Claim: 1
       9 Drawing Page(s)
DRWN
LN.CNT 1786
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
      The invention relates to improved methods of producing and recovering
AB
       sporulation-deficient B. anthracis mutant stains, and for
      producing and recovering recombinant B. anthracis protective
      antigen (PA), especially modified PA which is protease resistant, and to
      methods of using of these PAs or nucleic acids encoding these PAs for
      eliciting an immunogenic response in humans, including responses which
      provide protection against, or reduce the severity of, B.
      anthracis bacterial infections and which are useful to prevent
      and/or treat illnesses caused by B. anthracis, such as
       inhalation anthrax, cutaneous anthrax and
       gastrointestinal anthrax.
     ANSWER 10 OF 15 USPATFULL on STN
L5
AN
       2004:100777 USPATFULL
ΤI
       Methods for preparing bacillus anthracis protective antigen
       for use in vaccines
       Shiloach, Joseph, Rockville, MD, UNITED STATES
IN
         Leppla, Stephen H., Bethesda, MD, UNITED STATES
       Ramirez, Delia M., Bethesda, MD, UNITED STATES
       Schneerson, Rachel, Bethesda, MD, UNITED STATES
      Robbins, John B., Chevy Chase, MD, UNITED STATES
                               20040422
PΙ
      US 2004076638
                         A1
AΙ
      US 2002-290712
                         A1
                               20021108 (10)
                         20011109 (60)
PRAI
      US 2001-344505P
DT
      Utility
FS
      APPLICATION
      KLARQUIST SPARKMAN, LLP, 121 S.W. SALMON STREET, SUITE #1600, ONE WORLD
LREP
      TRADE CENTER, PORTLAND, OR, 97204-2988
CLMN
      Number of Claims: 35
ECL
      Exemplary Claim: 1
DRWN
      No Drawings
LN.CNT 1273
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
      The invention relates to improved methods of producing and recovering B.
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anthracisprotective antigen (PA), especially modified PA which is protease resistant, and to methods of using of these PAs or nucleic acids encoding these PAs for eliciting an immunogenic response in humans, including responses which provide protection against, or reduce the severity of, B. anthracis bacterial infections and which are useful to prevent and/or treat illnesses caused by B. anthracis, such as inhalation anthrax, cutaneous anthrax and gastrointestinal anthrax.

```
ANSWER 11 OF 15 USPATFULL on STN
L5
AN
       2003:140492 USPATFULL
       Anthrax lethal factor is a MAPK kinase protease
ΤI
IN
       Duesbery, Nicholas, Grand Rapids, MI, UNITED STATES
       Webb, Craig, Rockford, MI, UNITED STATES
         Leppla, Stephen, Bethesda, MD, UNITED STATES
       Vande Woude, George, Ada, MI, UNITED STATES
       US 2003096333
                         A1
                               20030522
PΙ
       US 6893835
                          B2
                               20050517
                               20020305 (10)
ΑI
       US 2002-93248
                          A1
RLI
       Division of Ser. No. US 2000-623104, filed on 13 Dec 2000, GRANTED, Pat.
       No. US 6485925
       Utility
DT
FS
       APPLICATION
       TOWNSEND AND TOWNSEND AND CREW, LLP, TWO EMBARCADERO CENTER, EIGHTH
LREP
       FLOOR, SAN FRANCISCO, CA, 94111-3834
CLMN
       Number of Claims: 61
ECL
       Exemplary Claim: 1
DRWN
       1 Drawing Page(s)
LN.CNT 2521
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
ΔR
       The present invention relates to in vitro and ex vivo methods of
       screening for modulators, homologues, and mimetics of lethal factor
       mitogen activated protein kinase kinase (MAPKK) protease activity, as
       well as methods of treating cancer by administering LF to transformed
       cells.
     ANSWER 12 OF 15 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on
L5
                                                        DUPLICATE 2
     2003:478120 BIOSIS
AN
     PREV200300478120
DN
     Poly(gamma-D-glutamic acid) protein conjugates induce IgG
TΙ
     antibodies in mice to the capsule of Bacillus anthracis: A
     potential addition to the anthrax vaccine.
     Schneerson, Rachel [Reprint Author]; Kubler-Kielb, Joanna; Liu, Teh-Yung;
ΑU
     Dai, Zhong-Dong; Leppla, Stephen H.; Yergey, Alfred; Backlund,
     Peter; Shiloach, Joseph; Majadly, Fathy; Robbins, John B.
CS
     National Institute of Child Health and Human Development, National
     Institutes of Health, Bethesda, MD, 20892, USA
     schneerr@mail.nih.gov
     Proceedings of the National Academy of Sciences of the United States of
SO
     America, (July 22 2003) Vol. 100, No. 15, pp. 8945-8950. print.
     ISSN: 0027-8424 (ISSN print).
DT
     Article
     English
LΑ
ED
     Entered STN: 15 Oct 2003
     Last Updated on STN: 15 Oct 2003
     Both the protective antigen (PA) and the poly(gamma-D-glutamic
     acid) capsule(gammaDPGA) are essential for the virulence of Bacillus
     anthracis. A critical level of vaccine-induced IgG anti-PA
     confers immunity to anthrax, but there is no information about
     the protective action of IgG anti-gammaDPGA. Because the number of spores
     presented by bioterrorists might be greater than encountered in nature, we
     sought to induce capsular antibodies to expand the immunity conferred by
     available anthrax vaccines. The nonimmunogenic gammaDPGA or
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corresponding synthetic peptides were bound to BSA, recombinant B. anthracis PA (rPA), or recombinant Pseudomonas aeruginosa exotoxin To identify the optimal construct, conjugates of B. anthracis gammaDPGA, Bacillus pumilus gammaDLPGA, and peptides of varying lengths (5-, 10-, or 20-mers), of the D or L configuration with active groups at the N or C termini, were bound at 5-32 mol per protein. The conjugates were characterized by physico-chemical and immunological assays, including GLC-MS and matrix-assisted laser desorption ionization time-of-flight spectrometry, and immunogenicity in 5- to 6-week-old mice. IqG anti-qammaDPGA and antiprotein were measured by ELISA. The highest levels of IgG anti-gammaDPGA were elicited by decamers of gammaDPGA at 10-20 mol per protein bound to the N- or C-terminal end. High IgG anti-gammaDPGA levels were elicited by two injections of 2.5 mug of levels of protein antibodies. rPA was the most effective carrier. Anti-gammaDPGA induced opsonophagocytic killing of B. anthracis tox-, cap+. gammaDPGA conjugates may enhance the protection conferred by PA alone. gammaDPGA-rPA conjugates induced both anti-PA and anti-gammaDPGA.

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ECL

1.5 AN

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IN

PA

PΙ

ΑI

PRAI

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gammaDPGA per mouse, whereas three injections were needed to achieve high
     ANSWER 13 OF 15 USPATFULL on STN
       2002:329841 USPATFULL
       Anthrax lethal factor is a MAPK kinase protease
       Duesbery, Nicholas, Grand Rapids, MI, UNITED STATES
       Webb, Craig, Rockford, MI, UNITED STATES
         Leppla, Stephen, Bethesda, MD, UNITED STATES
       Vande Woude, George, Ada, MI, UNITED STATES
       US 2002187521
                         A1
                               20021212
       US 6911203
                               20050628
                          B2
       US 2002-93200
                               20020305 (10)
                          A1
       Division of Ser. No. US 2000-623104, filed on 13 Dec 2000, ABANDONED A
       371 of International Ser. No. WO 1999-US7126, filed on 31 Mar 1999,
       PENDING
       US 1998-80330P
                           19980401 (60)
       Utility
       APPLICATION
       TOWNSEND AND TOWNSEND AND CREW, LLP, TWO EMBARCADERO CENTER, EIGHTH
       FLOOR, SAN FRANCISCO, CA, 94111-3834
       Number of Claims: 61
       Exemplary Claim: 1
       1 Drawing Page(s)
DRWN
LN.CNT 2519
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention relates to in vitro and ex vivo methods of
       screening for modulators, homologues, and mimetics of lethal factor
       mitogen activated protein kinase kinase (MAPKK) protease activity, as
       well as methods of treating cancer by administering LF to transformed
       cells.
     ANSWER 14 OF 15 USPATFULL on STN
       2002:310778 USPATFULL
       Anthrax lethal factor is a MAPK kinase protease
       Duesbery, Nicholas, Grand Rapids, MI, United States
       Webb, Craig, Rockford, MI, United States
         Leppla, Stephen, Bethesda, MD, United States
       Woude, George Vande, Ada, MI, United States
       The United States of America as represented by the Department of Health
       and Human Services, Washington, DC, United States (U.S. government)
       US 6485925
                          B1
                               20021126
       WO 9950439 19991007
       US 2000-623104
                               20001213 (9)
       WO 1999-US7126
                               19990331
                               20001213 PCT 371 date
      US 1998-80330P
                           19980401 (60)
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DT Utility FS GRANTED EXNAM Primary Examiner: Prouty, Rebecca E.; Assistant Examiner: Walicka, M. LREP Townsend and Townsend and Crew LLP CLMN Number of Claims: 15 ECL Exemplary Claim: 1 1 Drawing Figure(s); 1 Drawing Page(s) DRWN LN.CNT 2373 CAS INDEXING IS AVAILABLE FOR THIS PATENT. The present invention relates to in vitro and ex vivo methods of AB screening for modulators, homologues, and mimetics of lethal factor mitogen activated protein kinase kinase (MAPKK) protease activity, as well as methods of treating cancer by administering LF to transformed cells. ANSWER 15 OF 15 USPATFULL on STN 1.5 97:94207 USPATFULL AN TI Anthrax toxin fusion proteins and related methods Leppla, Stephen H., Bethesda, MD, United States IN Klimpel, Kurt R., Gaithersburg, MD, United States Arora, Naveen, Delhi, India Singh, Yogendra, Delhi, India Nichols, Peter J., Welling Kent, United Kingdom The Government of the United States as represented by the Secretary of PΑ the Department of Health and Human Services, Washington, DC, United States (U.S. government) PΙ US 5677274 19971014 ΑI US 1993-82849 19930625 (8) Continuation-in-part of Ser. No. US 1993-21601, filed on 12 Feb 1993, RLI now patented, Pat. No. US 5591631 DT Utility Granted FS EXNAM Primary Examiner: Jagannathan, Vasu S.; Assistant Examiner: Romeo, David Townsend and Townsend and Crew LREP Number of Claims: 12 CLMN Exemplary Claim: 1 ECL DRWN 1 Drawing Figure(s); 1 Drawing Page(s) LN.CNT 3382 CAS INDEXING IS AVAILABLE FOR THIS PATENT. AB The present invention provides a nucleic acid encoding a fusion protein comprising a nucleotide sequence encoding the anthrax protective antigen (PA) binding domain of the native anthrax lethal factor (LF) protein and a nucleotide sequence encoding an activity inducing domain of a second protein. Also provided is a nucleic acid encoding a fusion protein comprising a nucleotide sequence encoding the translocation domain and LF binding domain of the native anthrax PA protein and a nucleotide sequence encoding a ligand domain which specifically binds a cellular target. Proteins encoded by the nucleic acid of the invention, vectors comprising the nucleic acids and hosts capable of expressing the protein encoded by the nucleic acids are also provided. A composition comprising the PA binding domain of the native LF protein chemically attached to a non-LF activity inducing moiety is further provided. A method for delivering an activity to a cell is provided. The steps of the method include a) administering to the cell a protein comprising the translocation domain and the LF binding domain of the native PA protein and a ligand domain, and b) administering to the cell a product comprising the PA binding domain of the native LF protein and a non-LF activity inducing moiety, whereby the product administered in step b) is internalized into the cell and performs the activity within the cell. The invention also provides

proteins including an anthrax protective antigen which has been mutated to replace the trypsin cleavage site with residues

recognized specifically by the HIV-1 protease.

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=> e robbins john b/au
            87
                   ROBBINS JOHN/AU
E2
           126
                   ROBBINS JOHN A/AU
E3
           288 --> ROBBINS JOHN B/AU
                   ROBBINS JOHN C/AU
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                   ROBBINS JOHN CHAPMAN/AU
E5
E6
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            15
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E12
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=> d bib ab 1-
YOU HAVE REQUESTED DATA FROM 10 ANSWERS - CONTINUE? Y/(N):y
L7
     ANSWER 1 OF 10 USPATFULL on STN
       2006:158613 USPATFULL
AN
       Poly-gamma-glutamic conjugates for eliciting immune responses directed
ΤI
       against bacilli
       Schneerson, Rachel, Bethesda, MD, UNITED STATES
IN
       Leppla, Stephen, Bethesda, MD, UNITED STATES
         Robbins, John B., Chevy Chase, MD, UNITED STATES
       Shiloach, Joseph, Rockville, MD, UNITED STATES
       Kubler-Kielb, Joanna, Rockville, MD, UNITED STATES
       Liu, Darrell, Bethesda, MD, UNITED STATES
       Majadly, Fathy, Frederick, MD, UNITED STATES
PΙ
       US 2006134143
                          Αl
                               20060622
ΑI
       US 2004-559825
                          A1
                               20040604 (10)
       WO 2004-US17736
                               20040604
                                20051202 PCT 371 date
PRAI
       US 2003-476598P
                           20030605 (60)
DT
       Utility
FS
       APPLICATION
       KLARQUIST SPARKMAN, LLP, 121 S.W. SALMON STREET, SUITE #1600, PORTLAND,
LREP
       OR, 97204-2988, US
       Number of Claims: 36
CLMN
ECL
       Exemplary Claim: 1
DRWN
       2 Drawing Page(s)
LN.CNT 2866
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Immunogenic compositions and methods for eliciting an immune response
       against B. anthracis and other bacilli are provided that
       include immunogenic conjugates of a poly-γ-glutamic acid
       (\gamma PGA) polypeptide of B. anthracis, or of another
       Bacillus that expresses a \gammaPGA polypeptide. The \gammaPGA
       conjugates elicit an effective immune response against B.
       anthracis, or against another Bacillus, in mammalian hosts to
       which the conjugates are administered.
L7
     ANSWER 2 OF 10 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
     DUPLICATE 1
ΔN
     2006:436213 BIOSIS
     PREV200600430224
DM
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- TI Additional conjugation methods and immunogenicity of Bacillus anthracis poly-gamma-D-glutarnic acid-protein conjugates.
- AU Kubler-Kielb, Joanna [Reprint Author]; Liu, Teh-Yung; Mocca, Christopher; Majadly, Fathy; Robbins, John B.; Schneerson, Rachel
- CS NICHD, NIH, LDMI, 9000 Rockville Pike, Bldg 6, Rm 1A05, Bethesda, MD 20892 USA
 - kielbj@mail.nih.gov
- SO Infection and Immunity, (AUG 2006) Vol. 74, No. 8, pp. 4744-4749. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 30 Aug 2006
 - Last Updated on STN: 30 Aug 2006
- The capsule of Bacillus anthracis, composed of AB poly-gamma-D-glutamic acid (gamma DPGA), is an essential virulence factor of B. anthracis. The capsule inhibits innate host defense through its antiphagocytic action. gamma DPGA is a poor immunogen, but when covalently bound to a carrier protein, it elicits serum antibodies. To identify the optimal construct for clinical use, synthetic gamma DPGAs of different lengths were bound to carrier proteins at different densities. The advantages of the synthetic over the natural polypeptide are the homogeneous chain length and end groups, allowing conjugates to be accurately characterized and standardized and their chemical compositions to be related to their immunogenicities. In the present study, we evaluated, in addition to methods reported by us, hydrazone, oxime, and thioether linkages between gamma DPGA and several proteins, including bovine serum albumin, recombinant Pseudomonas aeruginosa exotoxin A, recombinant B. anthracis protective antigen (rPA), and tetanus toxoid (TT). The effects of the dosage and formulation on the immunogenicities of the conjugates were evaluated in mice. All conjugates were immunogenic. The optimal gamma DPGA chain length of 10 to 15 amino acids and the density, an average of 15 mol gamma DPGA per mol of protein, were confirmed. The thioether bond was the optimal linkage type, and TT and rPA were the best carriers. The optimal dosage was 1.2 to 2.5 mu g of gamma DPGA per mouse, and adsorption of the conjugates onto aluminum hydroxide significantly increased the antibody response to the protein with a lesser effect on anti-gamma DPGA levels.
- L7 ANSWER 3 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2
- AN 2005:1294042 CAPLUS
- DN 144:35295
- TI Hydrazone conjugates of haptens and antigens
- IN Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy; Leppla, Stephen H.; Robbins, John B.; Liu, Darrell T.; Shiloah, Joseph
- PA United States Dept. of Health and Human Services, USA
- SO U.S. Pat. Appl. Publ., 17 pp., Cont.-in-part of Appl. No. PCT/US04/017736. CODEN: USXXCO
- DT Patent
- LA English
- FAN.CNT 3

	PATENT NO.					D	DATE			APPL	ICAT	ION		DATE					
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ΡI	US 2005271675			A1 2005120			1208	US 2004-5851						20041206					
	WO 2005000884				A1 20050106			1	WO 2	004-1		20040604							
	WO 2005	C1 20051006																	
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     WO 2005117965
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              MR, NE, SN, TD, TG
PRAI WO 2004-US17736
                            A2
                                   20040604
     US 2003-476598P
                            P
                                   20030605
     US 2004-5851
                            A
                                   20041206
AB
     The authors disclose methods for making an immunogenic conjugate that
     includes a hapten or an antigen covalently linked to a carrier. The
     methods include reacting a first agent with a dihydrazide resulting in a
     hydrazino-modified first agent, wherein the first agent is a hapten, an
     antigen or a carrier; reacting a second agent with a benzaldehyde compound
     resulting in a benzaldehyde-modified second agent, wherein the second
     agent is a hapten, an antigen or a carrier, provided that the first agent
     or the second agent is a carrier; and reacting the hydrazine-modified
     first agent with the benzaldehyde-modified second agent resulting in an
     immunogenic conjugate comprising a hapten or an antigen covalently linked
     to a carrier via a hydrazone linkage.
     ANSWER 4 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN
L7
AN
     2005:1314018 CAPLUS
DN
     144:35300
     Methods for preparing immunogenic conjugates for use in vaccines
ΤI
IN
     Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy; Leppla, Stephen
     H.; Robbins, John B.; Liu, Darrell T.; Shiloach, Joseph
     The Government of the United States of America as Represented by the
PA
     Secretary, Department of Healthand Human Services, USA
SO
     PCT Int. Appl., 53 pp.
     CODEN: PIXXD2
DT
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LΆ
     English
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              LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
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                SN, TD, TG
      US 2005271675
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                                                                                     20041206
PRAI WO 2004-US17736
                                Α
                                        20040604
      US 2004-5851
                                Α
                                        20041206
      US 2003-476598P
                                Ρ
                                        20030605
      MARPAT 144:35300
OS
      Methods for making an immunogenic conjugate that includes a hapten or an
AB
      antigen covalently linked to a carrier are discussed. The methods include
      reacting a first agent with a dihydrazide resulting in a
      hydrazino-modified first agent, wherein the first agent is a hapten, an
      antigen or a carrier; reacting a second agent with a benzaldehyde compound
      resulting in a benzaldehyde-modified second agent, wherein the second
      agent is a hapten, an antigen or a carrier, provided that the first agent
      or the second agent is a carrier; and reacting the hydrazine-modified
      first agent with the benzaldehyde-modified second agent resulting in an
      immunogenic conjugate comprising a hapten or an antigen covalently linked
      to a carrier via a hydrazone linkage. The examples discuss the
      conjugation of Bacillus poly-γ-glutamic acids to carriers such as
      bovine serum albumin, Bacillus anthracis protective antigen, and
      Pseudomonas aeruginosa exotoxin A. The conjugates were shown to induce
      IgG and opsonophagocytosis.
                 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 4
                 ALL CITATIONS AVAILABLE IN THE RE FORMAT
L7
      ANSWER 5 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN
ΑN
      2005:14426 CAPLUS
DN
      142:112426
      Bacillus capsular poly-\gamma-glutamic acid conjugates for eliciting
TI
      immune responses against Bacillus infection
      Schneerson, Rachel; Leppla, Stephen; Robbins, John B.; Shiloach,
IN
      Joseph; Kubler-Kielb, Joanna; Liu, Darrell; Majadly, Fathy
      United States Dept. of Health and Human Services, USA
PA
SO
      PCT Int. Appl., 67 pp.
      CODEN: PIXXD2
DT
      Patent
LΑ
      English
FAN.CNT 3
      PATENT NO.
                               KIND
                                        DATE
                                                       APPLICATION NO.
                                                                                    DATE
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                                                       WO 2004-US17736
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ΡI
      WO 2005000884
                                A1
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                                20060622
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PRAI US 2003-476598P
                         P
                                20030605
     WO 2004-US17736
                        W
                                20040604
     US 2004-5851
                        A
                                20041206
     Immunogenic compns. and methods for eliciting an immune response against
AR
     B. anthracis and other bacilli are provided that include
     immunogenic conjugates of a poly-\gamma-glutamic acid (\gammaPGA)
     polypeptides of B. anthracis, or of another Bacillus that
     expresses a γPGA polypeptide. The γPGA conjugates elicit an
     effective immune response against B. anthracis, or against
     another Bacillus, in mammalian hosts to which the conjugates are
     administered. The conjugate consists of γ-D-PGA and carrier
     selected from bovine serum albumin, recombinant Bacillus protective
     antigen, recombinant Pseudomonas aeruginosa exotoxin A, tetanus toxoid,
     diphtheria toxoid, pertussis toxoid, Clostridium perfringens toxoid,
     HBsAg, HBcAg, heyhole limpet hemocyanin, horseshoe crab hemocyanin,
     edestin, mammalian serum albumin, mammalian Ig., analog or mimetic, or
     combination of two or more. The preferred conjugate consists of
     \gamma\text{-D-PGA} and Bacillus protective antigen.
RE.CNT 9
             THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
L7
     ANSWER 6 OF 10 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 3
     2004:331575 CAPLUS
AN
DN
     140:338027
TI
     Methods for preparing Bacillus anthracis protective antigen for
     use in vaccines
     Shiloach, Joseph; Leppla, Stephen H.; Ramirez, Delia M.; Schneerson,
IN
     Rachel; Robbins, John B.
PΑ
SO.
     U.S. Pat. Appl. Publ., 13 pp.
     CODEN: USXXCO
DT
     Patent
LA
     English
FAN.CNT 1
                                          APPLICATION NO.
     PATENT NO.
                        KIND
                               DATE
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                         A1
     US 2004076638
                               20040422
                                           US 2002-290712
                                                                   20021108
PRAI US 2001-344505P
                         P
                               20011109
     The authors disclose improved methods of producing and recovering B.
     anthracis protective antigen (PA), especially modified PA which is
     protease resistant, and to methods of using of these PAs or nucleic acids
     encoding these PAs for eliciting an immunogenic response.
L7
     ANSWER 7 OF 10 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
     2005:488548 BIOSIS
ΑN
     PREV200510291216
DN
TI
     Future vaccine development at NICHD.
     Robbins, John B. [Reprint Author]; Schneerson, Rachel
ΑU
     NICHD, Lab Dev and Mol Immun, Sect Bacterial Dis Pathogenesis and Immun,
CŞ
     NIH, Bldg 6, Room 436, Bethesda, MD 20892 USA
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robbinsj@nichd.nih.gov; schneerr@mail.nih.gov

SO Kaler, SG [Editor]; Rennert, OM [Editor]. (2004) pp. 49-59. Annals of the New York Academy of Sciences.

Publisher: NEW YORK ACAD SCIENCES, 2 EAST 63RD ST, NEW YORK, NY 10021 USA. Series: ANNALS OF THE NEW YORK ACADEMY OF SCIENCES.

Meeting Info.: 40th Scientific Symposium of the National-Institute-of-Child-Health-and-Human-Development. Bethesda, MD, USA. 20030908,. NICHD. ISSN: 0077-8923 (print). ISBN: 1-57331-520-6(H).

DT Book; (Book Chapter)
Conference; (Meeting)
General Review; (Literature Review)

LA English

ED Entered STN: 16 Nov 2005 Last Updated on STN: 16 Nov 2005

- Using published data and the results of our studies, we hypothesized that a critical level of serum IgG antibodies to the surface structures of invasive pathogens (capsular polysaccharides of Haemophilus influenzae type b, pneumococcus, meningococcus, Salmonella typhi, Escherichia coli, and Staphylococcus aureus, the O-specific polysaccharide LPS domain of the LPS of Shigella, non-typhoidal Salmonella, and E. coli, and the capsular polypeptide of Bacillus anthraces) confer immunity to these pathogens. Covalent attachment to a protein increases their immunogenicity and bestows T-cell properties to these antigens. We have also shown that a critical level of serum IgG antibodies to pertussis toxin alone induces immunity on both an individual and on a community basis (herd immunity) to Bordetella pertussis. It is likely that all the above conjugates and pertussis toxoid will be incorporated into vaccines for routine infant immunization.
- L7 ANSWER 8 OF 10 MEDLINE on STN
- AN 2005202078 MEDLINE
- DN PubMed ID: 15838097
- TI Future vaccine development at NICHD.
- AU Robbins John B; Schneerson Rachel
- CS Laboratory of Developmental and Molecular Immunity, NICHD, NIH, Building 6, Room 436, Bethesda, MD 20892, USA.. robbinsj@nichd.nih.gov
- SO Annals of the New York Academy of Sciences, (2004 Dec) Vol. 1038, pp. 49-59.

Journal code: 7506858. ISSN: 0077-8923.

- CY United States
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Priority Journals
- EM 200511
- ED Entered STN: 20 Apr 2005 Last Updated on STN: 10 Nov 2005 Entered Medline: 9 Nov 2005
- AB Using published data and the results of our studies, we hypothesized that a critical level of serum IgG antibodies to the surface structures of invasive pathogens (capsular polysaccharides of Haemophilus influenzae type b, pneumococcus, meningococcus, Salmonella typhi, Escherichia coli, and Staphylococcus aureus, the O-specific polysaccharide LPS domain of the LPS of Shigella, non-typhoidal Salmonella, and E. coli, and the capsular polypeptide of Bacillus anthraces) confer immunity to these pathogens. Covalent attachment to a protein increases their immunogenicity and bestows T-cell properties to these antigens. We have also shown that a critical level of serum IgG antibodies to pertussis toxin alone induces immunity on both an individual and on a community basis (herd immunity) to Bordetella pertussis. It is likely that all the above conjugates and pertussis toxoid will be incorporated into vaccines for routine infant immunization.
- L7 ANSWER 9 OF 10 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN DUPLICATE 4

- AN 2003:478120 BIOSIS
- DN PREV200300478120
- TI Poly(gamma-D-glutamic acid) protein conjugates induce IgG antibodies in mice to the capsule of Bacillus anthracis: A potential addition to the anthrax vaccine.
- AU Schneerson, Rachel [Reprint Author]; Kubler-Kielb, Joanna; Liu, Teh-Yung; Dai, Zhong-Dong; Leppla, Stephen H.; Yergey, Alfred; Backlund, Peter; Shiloach, Joseph; Majadly, Fathy; Robbins, John B.
- CS National Institute of Child Health and Human Development, National Institutes of Health, Bethesda, MD, 20892, USA schneerr@mail.nih.gov
- SO Proceedings of the National Academy of Sciences of the United States of America, (July 22 2003) Vol. 100, No. 15, pp. 8945-8950. print. ISSN: 0027-8424 (ISSN print).
- DT Article
- LA English
- ED Entered STN: 15 Oct 2003 Last Updated on STN: 15 Oct 2003
- AB Both the protective antigen (PA) and the poly(gamma-D-glutamic acid) capsule(gammaDPGA) are essential for the virulence of Bacillus anthracis. A critical level of vaccine-induced IgG anti-PA confers immunity to anthrax, but there is no information about the protective action of IgG anti-gammaDPGA. Because the number of spores presented by bioterrorists might be greater than encountered in nature, we sought to induce capsular antibodies to expand the immunity conferred by available anthrax vaccines. The nonimmunogenic gammaDPGA or corresponding synthetic peptides were bound to BSA, recombinant B. anthracis PA (rPA), or recombinant Pseudomonas aeruginosa exotoxin To identify the optimal construct, conjugates of B. A (rEPA). anthracis gammaDPGA, Bacillus pumilus gammaDLPGA, and peptides of varying lengths (5-, 10-, or 20-mers), of the D or L configuration with active groups at the N or C termini, were bound at 5-32 mol per protein. The conjugates were characterized by physico-chemical and immunological assays, including GLC-MS and matrix-assisted laser desorption ionization time-of-flight spectrometry, and immunogenicity in 5- to 6-week-old mice. IgG anti-gammaDPGA and antiprotein were measured by ELISA. The highest levels of IgG anti-gammaDPGA were elicited by decamers of gammaDPGA at 10-20 mol per protein bound to the N- or C-terminal end. High IgG anti-gammaDPGA levels were elicited by two injections of 2.5 mug of gammaDPGA per mouse, whereas three injections were needed to achieve high levels of protein antibodies. rPA was the most effective carrier. Anti-gammaDPGA induced opsonophagocytic killing of B. anthracis tox-, cap+. gammaDPGA conjugates may enhance the protection conferred by PA alone. gammaDPGA-rPA conjugates induced both anti-PA and anti-gammaDPGA.
- L7 ANSWER 10 OF 10 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN DUPLICATE 5
- AN 2002:429815 BIOSIS
- DN PREV200200429815
- TI Development of an improved vaccine for anthrax.
- AU Leppla, Stephen H. [Reprint author]; Robbins, John B.; Schneerson, Rachel; Shiloach, Joseph
- CS Oral Infection and Immunity Branch, NIDCR, 30 Convent Drive, Building 30, Room 303, Bethesda, MD, 20892-4350, USA
 Leppla@nih.gov
- SO Journal of Clinical Investigation, (July, 2002) Vol. 110, No. 2, pp. 141-144. print.

 CODEN: JCINAO. ISSN: 0021-9738.
- DT Article
- LA English
- ED Entered STN: 14 Aug 2002 Last Updated on STN: 14 Aug 2002

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L9
     ANSWER 1 OF 9 USPATFULL on STN
       2006:158613 USPATFULL
AN
       Poly-gamma-glutamic conjugates for eliciting immune responses directed
ΤТ
       against bacilli
       Schneerson, Rachel, Bethesda, MD, UNITED STATES
TN
       Leppla, Stephen, Bethesda, MD, UNITED STATES
       Robbins, John B., Chevy Chase, MD, UNITED STATES
         Shiloach, Joseph, Rockville, MD, UNITED STATES
       Kubler-Kielb, Joanna, Rockville, MD, UNITED STATES
       Liu, Darrell, Bethesda, MD, UNITED STATES
       Majadly, Fathy, Frederick, MD, UNITED STATES
PΙ
       US 2006134143
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AΙ
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                                20040604 (10)
       WO 2004-US17736
                                20040604
                                20051202 PCT 371 date
PRAI
       US 2003-476598P
                           20030605 (60)
DT
       Utility
FS
       APPLICATION
       KLARQUIST SPARKMAN, LLP, 121 S.W. SALMON STREET, SUITE #1600, PORTLAND,
LREP
       OR, 97204-2988, US
CLMN
       Number of Claims: 36
ECL
       Exemplary Claim: 1
DRWN
       2 Drawing Page(s)
LN.CNT 2866
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Immunogenic compositions and methods for eliciting an immune response
       against B. anthracis and other bacilli are provided that
       include immunogenic conjugates of a poly-\gamma-glutamic acid
       (\gamma PGA) polypeptide of B. anthracis, or of another
       Bacillus that expresses a \gammaPGA polypeptide. The \gammaPGA
       conjugates elicit an effective immune response against B.
       anthracis, or against another Bacillus, in mammalian hosts to
       which the conjugates are administered.
L9
     ANSWER 2 OF 9 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
     DUPLICATE 1
AN
     2006:459046 BIOSIS
DN
     PREV200600459246
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- TI Passive immunotherapy of Bacillus anthracis pulmonary infection in mice with antisera produced by DNA immunization.
- AU Herrmann, John E. [Reprint Author]; Wang, Shixia; Zhang, Chuanyou; Panchal, Rekha G.; Bavari, Sina; Lyons, C. Rick; Lovchik, Julie A.; Golding, Basil; Shiloach, Joseph; Lu, Shan
- CS Antibody Sci Inc, 80 Webster St, Worcester, MA 01603 USA ASI@AbScience.com
- SO Vaccine, (JUL 26 2006) Vol. 24, No. 31-32, pp. 5872-5880. CODEN: VACCDE. ISSN: 0264-410X.
- DT Article
- LA English
- ED Entered STN: 13 Sep 2006 Last Updated on STN: 13 Sep 2006
- Because of the high failure rate of antibiotic treatment in patients with AB anthrax there is a need for additional therapies such as passive immunization with therapeutic antibodies. In this study, we used codon-optimized plasmid DNAs (DNA vaccines) encoding Bacillus anthracis protective antigen (PA) to immunize rabbits for producing anti-anthrax antibodies for use in passive immunotherapy. The antisera generated with these DNA vaccines were of high titer as measured by ELISA. The antisera were also able to protect J774 macrophage cells by neutralizing the cytotoxic effect of exogenously added anthrax lethal toxin, and of the toxin released by B. anthracis (Sterne strain) spores following infection. addition, the antisera passively protected mice against pulmonary challenge with an approximate 50 LD50 dose of B. anthracis (Sterne strain) spores. The protection in mice was obtained when the antiserum was given 1 h before or 1 h after challenge. We further demonstrated that IgG and F(ab')(2) components purified from anti-PA rabbit hyperimmune sera retained similar levels of neutralizing activities against both exogenously added B. anthracis lethal toxin and toxin produced by B. anthracis (Sterne strain) spores. The high titer antisera we produced will enable an immunization strategy to supplement antibiotic therapy for improving the survival of patients with anthrax. (c) 2006 Elsevier Ltd. All rights reserved.
- L9 ANSWER 3 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2005:1314018 CAPLUS
- DN 144:35300
- TI Methods for preparing immunogenic conjugates for use in vaccines
- IN Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy; Leppla, Stephen H.; Robbins, John B.; Liu, Darrell T.; Shiloach, Joseph
- PA The Government of the United States of America as Represented by the Secretary, Department of Healthand Human Services, USA
- SO PCT Int. Appl., 53 pp.
- CODEN: PIXXD2
- DT Patent
- LA English
- FAN.CNT 3

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      US 2004-5851
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      US 2003-476598P
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os
      MARPAT 144:35300
AΒ
     Methods for making an immunogenic conjugate that includes a hapten or an
      antigen covalently linked to a carrier are discussed. The methods include
      reacting a first agent with a dihydrazide resulting in a
      hydrazino-modified first agent, wherein the first agent is a hapten, an
      antigen or a carrier; reacting a second agent with a benzaldehyde compound
      resulting in a benzaldehyde-modified second agent, wherein the second
      agent is a hapten, an antigen or a carrier, provided that the first agent
      or the second agent is a carrier; and reacting the hydrazine-modified
      first agent with the benzaldehyde-modified second agent resulting in an
      immunogenic conjugate comprising a hapten or an antigen covalently linked
      to a carrier via a hydrazone linkage. The examples discuss the
      conjugation of Bacillus poly-γ-glutamic acids to carriers such as
     bovine serum albumin, Bacillus anthracis protective antigen, and
      Pseudomonas aeruginosa exotoxin A. The conjugates were shown to induce
      IgG and opsonophagocytosis.
RE.CNT 4
                THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
                ALL CITATIONS AVAILABLE IN THE RE FORMAT
L9
      ANSWER 4 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
AN
      2005:14426 CAPLUS
DN
      142:112426
      Bacillus capsular poly-γ-glutamic acid conjugates for eliciting
ΤI
      immune responses against Bacillus infection
      Schneerson, Rachel; Leppla, Stephen; Robbins, John B.; Shiloach,
IN
      Joseph; Kubler-Kielb, Joanna; Liu, Darrell; Majadly, Fathy
PA
      United States Dept. of Health and Human Services, USA
so
      PCT Int. Appl., 67 pp.
      CODEN: PIXXD2
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     US 2006134143
                         A1
                                20060622
                                           US 2005-559825
                                                                  20051202
PRAI US 2003-476598P
                         P
                                20030605
     WO 2004-US17736
                         W
                                20040604
                                20041206
     US 2004-5851
                         Α
     Immunogenic compns. and methods for eliciting an immune response against
AΒ
     B. anthracis and other bacilli are provided that include
     immunogenic conjugates of a poly-γ-glutamic acid (γPGA)
     polypeptides of B. anthracis, or of another Bacillus that
     expresses a \gammaPGA polypeptide. The \gammaPGA conjugates elicit an
     effective immune response against B. anthracis, or against
     another Bacillus, in mammalian hosts to which the conjugates are
     administered. The conjugate consists of \gamma-D-PGA and carrier
     selected from bovine serum albumin, recombinant Bacillus protective
     antigen, recombinant Pseudomonas aeruginosa exotoxin A, tetanus toxoid,
     diphtheria toxoid, pertussis toxoid, Clostridium perfringens toxoid,
     HBsAq, HBcAq, heyhole limpet hemocyanin, horseshoe crab hemocyanin,
     edestin, mammalian serum albumin, mammalian Ig., analog or mimetic, or
     combination of two or more. The preferred conjugate consists of
     \gamma-D-PGA and Bacillus protective antigen.
RE.CNT 9
              THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
L9
     ANSWER 5 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2
AN
     2004:331575 CAPLUS
     140:338027
DN
     Methods for preparing Bacillus anthracis protective antigen for
TI
     use in vaccines
     Shiloach, Joseph; Leppla, Stephen H.; Ramirez, Delia M.;
IN
     Schneerson, Rachel; Robbins, John B.
PA
     U.S. Pat. Appl. Publ., 13 pp.
SO
     CODEN: USXXCO
DT
     Patent
LA
     English
FAN.CNT 1
     PATENT NO.
                        KIND
                                DATE
                                           APPLICATION NO.
                                                                   DATE
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                                           US 2002-290712
PΙ
     US 2004076638
                         A1
                                20040422
                                                                   20021108
PRAI US 2001-344505P
                         P
                                20011109
     The authors disclose improved methods of producing and recovering B.
     anthracis protective antigen (PA), especially modified PA which is
     protease resistant, and to methods of using of these PAs or nucleic acids
     encoding these PAs for eliciting an immunogenic response.
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ANSWER 6 OF 9 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN

L9

DUPLICATE 3

- AN 2004:149774 BIOSIS
- DN PREV200400153906
- TI Treatment of anthrax infection with combination of ciprofloxacin and antibodies to protective antigen of Bacillus anthracis.
- AU Karginov, Vladimir A. [Reprint Author]; Robinson, Tanisha M.; Riemenschneider, Jenny; Golding, Basil; Kennedy, Michael; Shiloach, Joseph; Alibek, Ken
- CS Advanced Biosystems, Inc., 10900 University Blvd., Manassas, VA, 20110, USA vladimir.karginov@analex.com
- SO FEMS Immunology and Medical Microbiology, (15 January 2004) Vol. 40, No. 1, pp. 71-74. print.
 ISSN: 0928-8244 (ISSN print).
- DT Article
- LA English
- ED Entered STN: 17 Mar 2004 Last Updated on STN: 17 Mar 2004
- Currently there is no effective treatment for inhalational anthrax beyond administration of antibiotics shortly after exposure. There is need for new, safe and effective treatments to supplement traditional antibiotic therapy. Our study was based on the premise that simultaneous inhibition of lethal toxin action with antibodies and blocking of bacterial growth by antibiotics will be beneficial for the treatment of anthrax. In this study, we tested the effects of a combination treatment using purified rabbit or sheep anti-protective antigen (PA) antibodies and the antibiotic ciprofloxacin in a rodent anthrax model. In mice infected with a dose of Bacillus anthracis Sterne strain corresponding to 10 LD50, antibiotic treatment with ciprofloxacin alone only cured 50% of infected animals. Administration of anti-PA IgG in combination with ciprofloxacin produced 90-100% survival. These data indicate that a combination of antibiotic/immunoglobulin therapy is more effective than antibiotic treatment alone in a rodent anthrax model.
- L9 ANSWER 7 OF 9 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN DUPLICATE 4
- AN 2003:478120 BIOSIS
- DN PREV200300478120
- TI Poly(gamma-D-glutamic acid) protein conjugates induce IgG antibodies in mice to the capsule of Bacillus anthracis: A potential addition to the anthrax vaccine.
- AU Schneerson, Rachel [Reprint Author]; Kubler-Kielb, Joanna; Liu, Teh-Yung; Dai, Zhong-Dong; Leppla, Stephen H.; Yergey, Alfred; Backlund, Peter; Shiloach, Joseph; Majadly, Fathy; Robbins, John B.
- CS National Institute of Child Health and Human Development, National Institutes of Health, Bethesda, MD, 20892, USA schneerr@mail.nih.gov
- SO Proceedings of the National Academy of Sciences of the United States of America, (July 22 2003) Vol. 100, No. 15, pp. 8945-8950. print. ISSN: 0027-8424 (ISSN print).
- DT Article
- LA English
- ED Entered STN: 15 Oct 2003 Last Updated on STN: 15 Oct 2003
- AB Both the protective antigen (PA) and the poly(gamma-D-glutamic acid) capsule(gammaDPGA) are essential for the virulence of Bacillus anthracis. A critical level of vaccine-induced IgG anti-PA confers immunity to anthrax, but there is no information about the protective action of IgG anti-gammaDPGA. Because the number of spores presented by bioterrorists might be greater than encountered in nature, we sought to induce capsular antibodies to expand the immunity conferred by available anthrax vaccines. The nonimmunogenic gammaDPGA or corresponding synthetic peptides were bound to BSA, recombinant B.

anthracis PA (rPA), or recombinant Pseudomonas aeruginosa exotoxin A (rEPA). To identify the optimal construct, conjugates of B. anthracis gammaDPGA, Bacillus pumilus gammaDLPGA, and peptides of varying lengths (5-, 10-, or 20-mers), of the D or L configuration with active groups at the N or C termini, were bound at 5-32 mol per protein. The conjugates were characterized by physico-chemical and immunological assays, including GLC-MS and matrix-assisted laser desorption ionization time-of-flight spectrometry, and immunogenicity in 5- to 6-week-old mice. IqG anti-qammaDPGA and antiprotein were measured by ELISA. The highest levels of IgG anti-gammaDPGA were elicited by decamers of gammaDPGA at 10-20 mol per protein bound to the N- or C-terminal end. High IgG anti-gammaDPGA levels were elicited by two injections of 2.5 mug of gammaDPGA per mouse, whereas three injections were needed to achieve high levels of protein antibodies. rPA was the most effective carrier. Anti-gammaDPGA induced opsonophagocytic killing of B. anthracis tox-, cap+. gammaDPGA conjugates may enhance the protection conferred by PA alone. gammaDPGA-rPA conjugates induced both anti-PA and anti-gammaDPGA.

- L9 ANSWER 8 OF 9 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN DUPLICATE 5
- AN 2002:281740 BIOSIS
- DN PREV200200281740
- TI Production, recovery and immunogenicity of the protective antigen from a recombinant strain of Bacillus anthracis.
- AU Ramirez, D. M.; Leppla, S. H.; Schneerson, R.; Shiloach, J. [Reprint author]
- CS Biotechnology Unit, LCDB, NIDDK, National Institutes of Health (NIH), Bethesda, MD, 20892, USA
- SO Journal of Industrial Microbiology and Biotechnology, (April, 2002) Vol. 28, No. 4, pp. 232-238. print. ISSN: 1367-5435.
- DT Article
- LA English
- ED Entered STN: 8 May 2002 Last Updated on STN: 8 May 2002
- AB The protective antigen (PA) is one of the three components of the anthrax toxin. It is a secreted nontoxic protein with a molecular weight of 83 kDa and is the major component of the currently licensed human vaccine for anthrax. Due to limitations found in the existing vaccine formulation, it has been proposed that genetically modified PA may be more effective as a vaccine. The expression and the stability of two recombinant PA (rPA) variants, PA-SNKE-DELTAFF-E308D and PA-N657A, were studied. These proteins were expressed in the nonsporogenic avirulent strain BH445. Initial results indicated that PA-SNKE-DELTAFF-E308D, which lacks two proteolysis-sensitive sites, is more stable than PA-N657A. Process development was conducted to establish an efficient production and purification process for PA-SNKE-DELTAFF-E308D. pH, media composition, growth strategy and protease inhibitors composition were analyzed. The production process chosen was based on batch growth of B. anthracis using tryptone and yeast extract as the only source of carbon, pH control at 7.5, and antifoam 289. Optimal harvest time was 14-18 h after inoculation, and EDTA (5 mM) was added upon harvest for proteolysis control. Recovery of the rPA was performed by expanded-bed adsorption (EBA) on a hydrophobic interaction chromatography (HIC) resin, eliminating the need for centrifugation, microfiltration and diafiltration. The EBA step was followed by ion exchange and gel filtration. rPA yields before and after purification were 130 and 90 mg/l, respectively. The purified rPA, without further treatment, treated with small amounts of formalin or adsorbed on alum, induced, high levels of IgG anti-PA with neutralization activities.

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2002:429815 BIOSIS
DN
     PREV200200429815
TΙ
     Development of an improved vaccine for anthrax.
ΑU
     Leppla, Stephen H. [Reprint author]; Robbins, John B.; Schneerson, Rachel;
     Shiloach, Joseph
     Oral Infection and Immunity Branch, NIDCR, 30 Convent Drive, Building 30,
CS
     Room 303, Bethesda, MD, 20892-4350, USA
     Leppla@nih.gov
SO
     Journal of Clinical Investigation, (July, 2002) Vol. 110, No. 2, pp.
     141-144. print.
     CODEN: JCINAO. ISSN: 0021-9738.
DΤ
     Article
LΑ
     English
ED
     Entered STN: 14 Aug 2002
     Last Updated on STN: 14 Aug 2002
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L11 ANSWER 1 OF 6 USPATFULL on STN
ΔN
       2006:158613 USPATFULL
TТ
       Poly-gamma-glutamic conjugates for eliciting immune responses directed
       against bacilli
       Schneerson, Rachel, Bethesda, MD, UNITED STATES
IN
       Leppla, Stephen, Bethesda, MD, UNITED STATES
       Robbins, John B., Chevy Chase, MD, UNITED STATES
       Shiloach, Joseph, Rockville, MD, UNITED STATES
         Kubler-Kielb, Joanna, Rockville, MD, UNITED STATES
       Liu, Darrell, Bethesda, MD, UNITED STATES
       Majadly, Fathy, Frederick, MD, UNITED STATES
PΤ
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       US 2004-559825
AΙ
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                                20040604 (10)
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                           20030605 (60)
DT
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FS
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LREP
       KLARQUIST SPARKMAN, LLP, 121 S.W. SALMON STREET, SUITE #1600, PORTLAND,
       OR, 97204-2988, US
CLMN
       Number of Claims: 36
ECL
       Exemplary Claim: 1
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ΔN

DRWN 2 Drawing Page(s)

LN.CNT 2866

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Immunogenic compositions and methods for eliciting an immune response against B. anthracis and other bacilli are provided that include immunogenic conjugates of a poly-γ-glutamic acid (γPGA) polypeptide of B. anthracis, or of another Bacillus that expresses a γPGA polypeptide. The γPGA conjugates elicit an effective immune response against B. anthracis, or against another Bacillus, in mammalian hosts to which the conjugates are administered.

- L11 ANSWER 2 OF 6 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN DUPLICATE 1
- AN 2006:436213 BIOSIS
- DN PREV200600430224
- TI Additional conjugation methods and immunogenicity of Bacillus anthracis poly-gamma-D-glutarnic acid-protein conjugates.
- AU Kubler-Kielb, Joanna [Reprint Author]; Liu, Teh-Yung; Mocca, Christopher; Majadly, Fathy; Robbins, John B.; Schneerson, Rachel
- CS NICHD, NIH, LDMI, 9000 Rockville Pike, Bldg 6, Rm 1A05, Bethesda, MD 20892 USA

kielbj@mail.nih.gov

- SO Infection and Immunity, (AUG 2006) Vol. 74, No. 8, pp. 4744-4749. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 30 Aug 2006 Last Updated on STN: 30 Aug 2006
- AB The capsule of Bacillus anthracis, composed of poly-gamma-D-glutamic acid (gamma DPGA), is an essential virulence factor of B. anthracis. The capsule inhibits innate host defense through its antiphagocytic action. gamma DPGA is a poor immunogen, but when covalently bound to a carrier protein, it elicits serum antibodies. To identify the optimal construct for clinical use, synthetic gamma DPGAs of different lengths were bound to carrier proteins at different densities. The advantages of the synthetic over the natural polypeptide are the homogeneous chain length and end groups, allowing conjugates to be accurately characterized and standardized and their chemical compositions to be related to their immunogenicities. In the present study, we evaluated, in addition to methods reported by us, hydrazone, oxime, and thioether linkages between gamma DPGA and several proteins, including bovine serum albumin, recombinant Pseudomonas aeruginosa exotoxin A, recombinant B. anthracis protective antigen (rPA), and tetanus toxoid (TT). The effects of the dosage and formulation on the immunogenicities of the conjugates were evaluated in mice. All conjugates were immunogenic. The optimal gamma DPGA chain length of 10 to 15 amino acids and the density, an average of 15 mol gamma DPGA per mol of protein, were confirmed. The thioether bond was the optimal linkage type, and TT and rPA were the best carriers. The optimal dosage was 1.2 to 2.5 mu g of gamma DPGA per mouse, and adsorption of the conjugates onto aluminum hydroxide significantly increased the antibody response to the protein with a lesser effect on anti-gamma DPGA levels.
- L11 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2
- AN 2005:1294042 CAPLUS
- DN 144:35295
- TI Hydrazone conjugates of haptens and antigens
- IN Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy;
 Leppla, Stephen H.; Robbins, John B.; Liu, Darrell T.; Shiloah, Joseph
- PA United States Dept. of Health and Human Services, USA
- SO U.S. Pat. Appl. Publ., 17 pp., Cont.-in-part of Appl. No. PCT/US04/017736. CODEN: USXXCO
- DT Patent

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     WO 2005000884
                         A1
                               20050106
     WO 2005000884
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                               20051006
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PRAI WO 2004-US17736
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                               20040604
                               20030605
     US 2003-476598P
                         Ρ
     US 2004-5851
                         Α
                               20041206
     The authors disclose methods for making an immunogenic conjugate that
AΒ
     includes a hapten or an antigen covalently linked to a carrier. The
     methods include reacting a first agent with a dihydrazide resulting in a
     hydrazino-modified first agent, wherein the first agent is a hapten, an
     antigen or a carrier; reacting a second agent with a benzaldehyde compound
     resulting in a benzaldehyde-modified second agent, wherein the second
     agent is a hapten, an antigen or a carrier, provided that the first agent
     or the second agent is a carrier; and reacting the hydrazine-modified
     first agent with the benzaldehyde-modified second agent resulting in an
     immunogenic conjugate comprising a hapten or an antigen covalently linked
     to a carrier via a hydrazone linkage.
L11
    ANSWER 4 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN
     2005:1314018 CAPLUS
AN
DN
     144:35300
    Methods for preparing immunogenic conjugates for use in vaccines
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IN
     Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy;
     Leppla, Stephen H.; Robbins, John B.; Liu, Darrell T.; Shiloach, Joseph
     The Government of the United States of America as Represented by the
PA
     Secretary, Department of Healthand Human Services, USA
so
     PCT Int. Appl., 53 pp.
    CODEN: PIXXD2
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    English
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     MARPAT 144:35300
os
    Methods for making an immunogenic conjugate that includes a hapten or an
AΒ
     antigen covalently linked to a carrier are discussed. The methods include
     reacting a first agent with a dihydrazide resulting in a
     hydrazino-modified first agent, wherein the first agent is a hapten, an
     antigen or a carrier; reacting a second agent with a benzaldehyde compound
     resulting in a benzaldehyde-modified second agent, wherein the second
     agent is a hapten, an antigen or a carrier, provided that the first agent
     or the second agent is a carrier; and reacting the hydrazine-modified
     first agent with the benzaldehyde-modified second agent resulting in an
     immunogenic conjugate comprising a hapten or an antigen covalently linked
     to a carrier via a hydrazone linkage. The examples discuss the
     conjugation of Bacillus poly-γ-glutamic acids to carriers such as
     bovine serum albumin, Bacillus anthracis protective antigen, and
     Pseudomonas aeruginosa exotoxin A. The conjugates were shown to induce
     IqG and opsonophagocytosis.
             THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
    ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN
L11
                CAPLUS
AN
     2005:14426
DN
     142:112426
ΤI
     Bacillus capsular poly-γ-glutamic acid conjugates for eliciting
     immune responses against Bacillus infection
IN
     Schneerson, Rachel; Leppla, Stephen; Robbins, John B.; Shiloach, Joseph;
     Kubler-Kielb, Joanna; Liu, Darrell; Majadly, Fathy
PA
     United States Dept. of Health and Human Services, USA
SO
     PCT Int. Appl., 67 pp.
     CODEN: PIXXD2
DT
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     B. anthracis and other bacilli are provided that include
     immunogenic conjugates of a poly-\gamma-glutamic acid (\gammaPGA)
     polypeptides of B. anthracis, or of another Bacillus that
     expresses a \gammaPGA polypeptide. The \gammaPGA conjugates elicit an
     effective immune response against B. anthracis, or against
     another Bacillus, in mammalian hosts to which the conjugates are
                   The conjugate consists of \gamma-D-PGA and carrier
     administered.
     selected from bovine serum albumin, recombinant Bacillus protective
     antigen, recombinant Pseudomonas aeruginosa exotoxin A, tetanus toxoid,
     diphtheria toxoid, pertussis toxoid, Clostridium perfringens toxoid,
     HBsAg, HBcAg, heyhole limpet hemocyanin, horseshoe crab hemocyanin,
     edestin, mammalian serum albumin, mammalian Ig., analog or mimetic, or
     combination of two or more. The preferred conjugate consists of
     \gamma-D-PGA and Bacillus protective antigen.
RE.CNT 9
              THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
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- L11 ANSWER 6 OF 6 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN DUPLICATE 3
- 2003:478120 BIOSIS AN
- PREV200300478120 DN
- ΤI Poly(gamma-D-glutamic acid) protein conjugates induce IgG antibodies in mice to the capsule of Bacillus anthracis: A potential addition to the anthrax vaccine.
- ΑU Schneerson, Rachel [Reprint Author]; Kubler-Kielb, Joanna; Liu, Teh-Yung; Dai, Zhong-Dong; Leppla, Stephen H.; Yergey, Alfred; Backlund, Peter; Shiloach, Joseph; Majadly, Fathy; Robbins, John B.
- National Institute of Child Health and Human Development, National CS Institutes of Health, Bethesda, MD, 20892, USA schneerr@mail.nih.gov

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SO Proceedings of the National Academy of Sciences of the United States of America, (July 22 2003) Vol. 100, No. 15, pp. 8945-8950. print. ISSN: 0027-8424 (ISSN print).
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DT Article

LA English

ED Entered STN: 15 Oct 2003 Last Updated on STN: 15 Oct 2003

Both the protective antigen (PA) and the poly(gamma-D-glutamic acid) AB capsule (qammaDPGA) are essential for the virulence of Bacillus anthracis. A critical level of vaccine-induced IqG anti-PA confers immunity to anthrax, but there is no information about the protective action of IgG anti-gammaDPGA. Because the number of spores presented by bioterrorists might be greater than encountered in nature, we sought to induce capsular antibodies to expand the immunity conferred by available anthrax vaccines. The nonimmunogenic gammaDPGA or corresponding synthetic peptides were bound to BSA, recombinant B. anthracis PA (rPA), or recombinant Pseudomonas aeruginosa exotoxin A (rEPA). To identify the optimal construct, conjugates of B. anthracis gammaDPGA, Bacillus pumilus gammaDLPGA, and peptides of varying lengths (5-, 10-, or 20-mers), of the D or L configuration with active groups at the N or C termini, were bound at 5-32 mol per protein. The conjugates were characterized by physico-chemical and immunological assays, including GLC-MS and matrix-assisted laser desorption ionization time-of-flight spectrometry, and immunogenicity in 5- to 6-week-old mice. IgG anti-gammaDPGA and antiprotein were measured by ELISA. The highest levels of IgG anti-gammaDPGA were elicited by decamers of gammaDPGA at 10-20 mol per protein bound to the N- or C-terminal end. High IgG anti-gammaDPGA levels were elicited by two injections of 2.5 mug of gammaDPGA per mouse, whereas three injections were needed to achieve high levels of protein antibodies. rPA was the most effective carrier. Anti-gammaDPGA induced opsonophagocytic killing of B. anthracis tox-, cap+. gammaDPGA conjugates may enhance the protection conferred by PA alone. gammaDPGA-rPA conjugates induced both anti-PA and anti-gammaDPGA.

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L13 ANSWER 1 OF 4 USPATFULL on STN
AN
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Poly-gamma-glutamic conjugates for eliciting immune responses directed
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         against bacilli
         Schneerson, Rachel, Bethesda, MD, UNITED STATES
IN
        Leppla, Stephen, Bethesda, MD, UNITED STATES
        Robbins, John B., Chevy Chase, MD, UNITED STATES
        Shiloach, Joseph, Rockville, MD, UNITED STATES
        Kubler-Kielb, Joanna, Rockville, MD, UNITED STATES
           Liu, Darrell, Bethesda, MD, UNITED STATES
        Majadly, Fathy, Frederick, MD, UNITED STATES
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        KLARQUIST SPARKMAN, LLP, 121 S.W. SALMON STREET, SUITE #1600, PORTLAND,
LREP
        OR, 97204-2988, US
CLMN
        Number of Claims: 36
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        Exemplary Claim: 1
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         Immunogenic compositions and methods for eliciting an immune response
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        against B. anthracis and other bacilli are provided that
         include immunogenic conjugates of a poly-γ-glutamic acid
         (\gamma PGA) polypeptide of B. anthracis, or of another
        Bacillus that expresses a \gammaPGA polypeptide. The \gammaPGA
        conjugates elicit an effective immune response against B.
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      ANSWER 2 OF 4 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 1
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AN
      2005:1294042 CAPLUS
DN
      144:35295
      Hydrazone conjugates of haptens and antigens
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      Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy; Leppla, Stephen
IN
      H.; Robbins, John B.; Liu, Darrell T.; Shiloah, Joseph
      United States Dept. of Health and Human Services, USA
PA
      U.S. Pat. Appl. Publ., 17 pp., Cont.-in-part of Appl. No. PCT/US04/017736.
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                                   20030605
     US 2004-5851
                            Α
                                   20041206
     The authors disclose methods for making an immunogenic conjugate that
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     includes a hapten or an antigen covalently linked to a carrier.
     methods include reacting a first agent with a dihydrazide resulting in a
     hydrazino-modified first agent, wherein the first agent is a hapten, an
     antigen or a carrier; reacting a second agent with a benzaldehyde compound
     resulting in a benzaldehyde-modified second agent, wherein the second
     agent is a hapten, an antigen or a carrier, provided that the first agent
     or the second agent is a carrier; and reacting the hydrazine-modified
     first agent with the benzaldehyde-modified second agent resulting in an
     immunogenic conjugate comprising a hapten or an antigen covalently linked
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                             COPYRIGHT 2006 ACS on STN
     ANSWER 3 OF 4 CAPLUS
L13
     2005:1314018 CAPLUS
AN
DN
     144:35300
ΤI
     Methods for preparing immunogenic conjugates for use in vaccines
     Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy; Leppla, Stephen
IN
     H.; Robbins, John B.; Liu, Darrell T.; Shiloach, Joseph
     The Government of the United States of America as Represented by the
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     Secretary, Department of Healthand Human Services, USA
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AB
    Methods for making an immunogenic conjugate that includes a hapten or an
     antigen covalently linked to a carrier are discussed. The methods include
     reacting a first agent with a dihydrazide resulting in a
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hydrazino-modified first agent, wherein the first agent is a hapten, an antigen or a carrier; reacting a second agent with a benzaldehyde compound resulting in a benzaldehyde-modified second agent, wherein the second agent is a hapten, an antigen or a carrier, provided that the first agent or the second agent is a carrier; and reacting the hydrazine-modified first agent with the benzaldehyde-modified second agent resulting in an immunogenic conjugate comprising a hapten or an antigen covalently linked to a carrier via a hydrazone linkage. The examples discuss the conjugation of Bacillus poly-γ-qlutamic acids to carriers such as bovine serum albumin, Bacillus anthracis protective antigen, and Pseudomonas aeruginosa exotoxin A. The conjugates were shown to induce IgG and opsonophagocytosis.

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

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ANSWER 4 OF 4 CAPLUS COPYRIGHT 2006 ACS on STN
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2005:14426 CAPLUS AN

DN

Bacillus capsular poly-γ-glutamic acid conjugates for eliciting TIimmune responses against Bacillus infection

IN Schneerson, Rachel; Leppla, Stephen; Robbins, John B.; Shiloach, Joseph; Kubler-Kielb, Joanna; Liu, Darrell; Majadly, Fathy

United States Dept. of Health and Human Services, USA PA

SO PCT Int. Appl., 67 pp. CODEN: PIXXD2

Patent

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     Immunogenic compns. and methods for eliciting an immune response against
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     B. anthracis and other bacilli are provided that include
     immunogenic conjugates of a poly-\gamma-glutamic acid (\gammaPGA)
     polypeptides of B. anthracis, or of another Bacillus that
     expresses a \gamma PGA polypeptide. The \gamma PGA conjugates elicit an
     effective immune response against B. anthracis, or against
     another Bacillus, in mammalian hosts to which the conjugates are
     administered. The conjugate consists of \gamma-D-PGA and carrier
     selected from bovine serum albumin, recombinant Bacillus protective
     antigen, recombinant Pseudomonas aeruginosa exotoxin A, tetanus toxoid,
     diphtheria toxoid, pertussis toxoid, Clostridium perfringens toxoid,
     HBsAg, HBcAg, heyhole limpet hemocyanin, horseshoe crab hemocyanin,
     edestin, mammalian serum albumin, mammalian Ig., analog or mimetic, or
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     \gamma-D-PGA and Bacillus protective antigen.
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       against B. anthracis and other bacilli are provided that
       include immunogenic conjugates of a poly-γ-glutamic acid
       (\gamma PGA) polypeptide of B. anthracis, or of another
       Bacillus that expresses a \gamma PGA polypeptide. The \gamma PGA
       conjugates elicit an effective immune response against B.
       anthracis, or against another Bacillus, in mammalian hosts to
       which the conjugates are administered.
L15
    ANSWER 2 OF 6 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
     DUPLICATE 1
     2006:436213 BIOSIS
ΔN
DN
     PREV200600430224
TТ
     Additional conjugation methods and immunogenicity of Bacillus
     anthracis poly-gamma-D-glutarnic acid-protein conjugates.
ΔII
     Kubler-Kielb, Joanna [Reprint Author]; Liu, Teh-Yung; Mocca, Christopher;
     Majadly, Fathy; Robbins, John B.; Schneerson, Rachel
CS
     NICHD, NIH, LDMI, 9000 Rockville Pike, Bldg 6, Rm 1A05, Bethesda, MD 20892
     USA
     kielbj@mail.nih.gov
SO
     Infection and Immunity, (AUG 2006) Vol. 74, No. 8, pp. 4744-4749.
     CODEN: INFIBR. ISSN: 0019-9567.
DT
     Article
     English
LA
     Entered STN: 30 Aug 2006
ED
     Last Updated on STN: 30 Aug 2006
AB
     The capsule of Bacillus anthracis, composed of
     poly-gamma-D-glutamic acid (gamma DPGA), is an essential virulence factor
     of B. anthracis. The capsule inhibits innate host defense
     through its antiphagocytic action. gamma DPGA is a poor immunogen, but
     when covalently bound to a carrier protein, it elicits serum antibodies.
     To identify the optimal construct for clinical use, synthetic gamma DPGAs
     of different lengths were bound to carrier proteins at different
     densities. The advantages of the synthetic over the natural polypeptide
     are the homogeneous chain length and end groups, allowing conjugates to be
     accurately characterized and standardized and their chemical compositions
     to be related to their immunogenicities. In the present study, we
     evaluated, in addition to methods reported by us, hydrazone, oxime, and
     thioether linkages between gamma DPGA and several proteins, including
     bovine serum albumin, recombinant Pseudomonas aeruginosa exotoxin A,
     recombinant B. anthracis protective antigen (rPA), and tetanus
     toxoid (TT). The effects of the dosage and formulation on the
     immunogenicities of the conjugates were evaluated in mice. All conjugates
     were immunogenic. The optimal gamma DPGA chain length of 10 to 15 amino
     acids and the density, an average of 15 mol gamma DPGA per mol of protein,
     were confirmed. The thioether bond was the optimal linkage type, and TT
     and rPA were the best carriers. The optimal dosage was 1.2 to 2.5 mu g of
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gamma DPGA per mouse, and adsorption of the conjugates onto aluminum

hydroxide significantly increased the antibody response to the protein with a lesser effect on anti-gamma DPGA levels.

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L15
     ANSWER 3 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2
     2005:1294042 CAPLUS
AN
DN
     144:35295
ΤI
     Hydrazone conjugates of haptens and antigens
IN
     Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy;
     Leppla, Stephen H.; Robbins, John B.; Liu, Darrell T.; Shiloah, Joseph
PA
     United States Dept. of Health and Human Services, USA
so
     U.S. Pat. Appl. Publ., 17 pp., Cont.-in-part of Appl. No. PCT/US04/017736.
     CODEN: USXXCO
DT
     Patent
LA
     English
FAN.CNT 3
     PATENT NO.
                        KIND
                                DATE
                                           APPLICATION NO.
                                                                   DATE
                                            ______
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     US 2005271675
                         A1
                                20051208
                                            US 2004-5851
                                                                   20041206
     WO 2005000884
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     WO 2005117965
                          A1
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             RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
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PRAI WO 2004-US17736
                         A2
                                20040604
     US 2003-476598P
                          P
                                20030605
     US 2004-5851
                          Α
                                20041206
AB
     The authors disclose methods for making an immunogenic conjugate that
     includes a hapten or an antigen covalently linked to a carrier. The
     methods include reacting a first agent with a dihydrazide resulting in a
     hydrazino-modified first agent, wherein the first agent is a hapten, an
     antigen or a carrier; reacting a second agent with a benzaldehyde compound
```

to a carrier via a hydrazone linkage.

resulting in a benzaldehyde-modified second agent, wherein the second agent is a hapten, an antigen or a carrier, provided that the first agent or the second agent is a carrier; and reacting the hydrazine-modified first agent with the benzaldehyde-modified second agent resulting in an immunogenic conjugate comprising a hapten or an antigen covalently linked

L15 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

AN 2005:1314018 CAPLUS

DN 144:35300

TI Methods for preparing immunogenic conjugates for use in vaccines

IN Schneerson, Rachel; Kubler-Kielb, Joanna; Majadly, Fathy;

Leppla, Stephen H.; Robbins, John B.; Liu, Darrell T.; Shiloach, Joseph The Government of the United States of America as Represented by the PA Secretary, Department of Healthand Human Services, USA SO PCT Int. Appl., 53 pp. CODEN: PIXXD2 DT Patent T.A English FAN.CNT 3 PATENT NO. KIND DATE APPLICATION NO. ---------20051215 WO 2005-US19678 WO 2005117965 A1 20050603 PΙ AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG WO 2004-US17736 WO 2005000884 20050106 20040604 A1 WO 2005000884 C1 20051006 AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, W: CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG 20051208 US 2004-5851 20041206 US 2005271675 A1 PRAI WO 2004-US17736 Α 20040604 US 2004-5851 Α 20041206 US 2003-476598P P 20030605 MARPAT 144:35300 os Methods for making an immunogenic conjugate that includes a hapten or an AB antigen covalently linked to a carrier are discussed. The methods include reacting a first agent with a dihydrazide resulting in a hydrazino-modified first agent, wherein the first agent is a hapten, an antigen or a carrier; reacting a second agent with a benzaldehyde compound resulting in a benzaldehyde-modified second agent, wherein the second agent is a hapten, an antigen or a carrier, provided that the first agent or the second agent is a carrier; and reacting the hydrazine-modified first agent with the benzaldehyde-modified second agent resulting in an immunogenic conjugate comprising a hapten or an antigen covalently linked to a carrier via a hydrazone linkage. The examples discuss the conjugation of Bacillus poly-γ-glutamic acids to carriers such as bovine serum albumin, Bacillus anthracis protective antigen, and Pseudomonas aeruginosa exotoxin A. The conjugates were shown to induce IgG and opsonophagocytosis. RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

AN 2005:14426 CAPLUS DN 142:112426 TI Bacillus capsular poly- γ -glutamic acid conjugates for eliciting immune responses against Bacillus infection

ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

L15

IN Schneerson, Rachel; Leppla, Stephen; Robbins, John B.; Shiloach, Joseph; Kubler-Kielb, Joanna; Liu, Darrell; Majadly, Fathy PA United States Dept. of Health and Human Services, USA PCT Int. Appl., 67 pp. SO CODEN: PIXXD2 DT Patent English LA FAN.CNT 3 PATENT NO. KIND DATE APPLICATION NO. DATE ------_ _ _ _ ----------_____ WO 2005000884 WO 2004-US17736 ΡI **A**1 20050106 20040604 WO 2005000884 C1 20051006 AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG AU 2004252091 20050106 AU 2004-252091 **A1** 20040604 CA 2528067 AA 20050106 CA 2004-2528067 20040604 EP 2004-754360 EP 1633778 **A1** 20060315 20040604 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK 20051208 US 2004-5851 US 2005271675 A1 20041206 WO 2005117965 **A1** 20051215 WO 2005-US19678 20050603 AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, W: CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG US 2005-559825 20051202 US 2006134143 A1 20060622 PRAI US 2003-476598P Ρ 20030605 WO 2004-US17736 W 20040604 US 2004-5851 Α 20041206 Immunogenic compns. and methods for eliciting an immune response against AΒ B. anthracis and other bacilli are provided that include immunogenic conjugates of a poly-γ-glutamic acid (γPGA) polypeptides of B. anthracis, or of another Bacillus that expresses a γ PGA polypeptide. The γ PGA conjugates elicit an effective immune response against B. anthracis, or against another Bacillus, in mammalian hosts to which the conjugates are administered. The conjugate consists of γ -D-PGA and carrier selected from bovine serum albumin, recombinant Bacillus protective antigen, recombinant Pseudomonas aeruginosa exotoxin A, tetanus toxoid, diphtheria toxoid, pertussis toxoid, Clostridium perfringens toxoid, HBsAg, HBcAg, heyhole limpet hemocyanin, horseshoe crab hemocyanin, edestin, mammalian serum albumin, mammalian Ig., analog or mimetic, or combination of two or more. The preferred conjugate consists of γ -D-PGA and Bacillus protective antigen.

RE.CNT 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L15 ANSWER 6 OF 6 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN DUPLICATE 3
- AN 2003:478120 BIOSIS
- DN PREV200300478120
- Poly(gamma-D-glutamic acid) protein conjugates induce IgG antibodies in mice to the capsule of Bacillus anthracis: A potential addition to the anthrax vaccine.
- AU Schneerson, Rachel [Reprint Author]; Kubler-Kielb, Joanna; Liu, Teh-Yung; Dai, Zhong-Dong; Leppla, Stephen H.; Yergey, Alfred; Backlund, Peter; Shiloach, Joseph; Majadly, Fathy; Robbins, John B.
- CS National Institute of Child Health and Human Development, National Institutes of Health, Bethesda, MD, 20892, USA schneerr@mail.nih.gov
- SO Proceedings of the National Academy of Sciences of the United States of America, (July 22 2003) Vol. 100, No. 15, pp. 8945-8950. print. ISSN: 0027-8424 (ISSN print).
- DT Article
- LA English
- ED Entered STN: 15 Oct 2003 Last Updated on STN: 15 Oct 2003
- Both the protective antigen (PA) and the poly(gamma-D-glutamic acid) ΔR capsule (qammaDPGA) are essential for the virulence of Bacillus anthracis. A critical level of vaccine-induced IgG anti-PA confers immunity to anthrax, but there is no information about the protective action of IgG anti-gammaDPGA. Because the number of spores presented by bioterrorists might be greater than encountered in nature, we sought to induce capsular antibodies to expand the immunity conferred by available anthrax vaccines. The nonimmunogenic gammaDPGA or corresponding synthetic peptides were bound to BSA, recombinant B. anthracis PA (rPA), or recombinant Pseudomonas aeruginosa exotoxin A (rEPA). To identify the optimal construct, conjugates of B. anthracis gammaDPGA, Bacillus pumilus gammaDLPGA, and peptides of varying lengths (5-, 10-, or 20-mers), of the D or L configuration with active groups at the N or C termini, were bound at 5-32 mol per protein. The conjugates were characterized by physico-chemical and immunological assays, including GLC-MS and matrix-assisted laser desorption ionization time-of-flight spectrometry, and immunogenicity in 5- to 6-week-old mice. IgG anti-gammaDPGA and antiprotein were measured by ELISA. The highest levels of IgG anti-gammaDPGA were elicited by decamers of gammaDPGA at 10-20 mol per protein bound to the N- or C-terminal end. High IgG anti-gammaDPGA levels were elicited by two injections of 2.5 mug of gammaDPGA per mouse, whereas three injections were needed to achieve high levels of protein antibodies. rPA was the most effective carrier. Anti-gammaDPGA induced opsonophagocytic killing of B. anthracis tox-, cap+. gammaDPGA conjugates may enhance the protection conferred by PA alone. gammaDPGA-rPA conjugates induced both anti-PA and anti-gammaDPGA.

=> s conjugate? and ?PGA
'?PGA' NOT LONG ENOUGH FOR LEFT TRUNCATION
You have entered a truncated stem whose length is less than the minimum allowed for left truncation in the requested search field. You may increase the length of the stem to the minimum allowed and try again. Enter HELP SFIELDS to to find the minimum stem length for left truncation in the requested search field.

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PROCESSING COMPLETED FOR L16
L17 22 DUP REM L16 (0 DUPLICATES REMOVED)

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=> d bib ab 1-
YOU HAVE REQUESTED DATA FROM 22 ANSWERS - CONTINUE? Y/(N):y
     ANSWER 1 OF 22 USPATFULL on STN
AN
       2006:166354 USPATFULL
       Drug pre-targeting by means of bi-specific antibodies and hapten
TI
       constructs comprising a carrier peptide and the active agent(s)
IN
       Goldenberg, David M, Mendham, NJ, UNITED STATES
       Hansen, Hans J, Picayune, NJ, UNITED STATES
       Leung, Shui-on, Hong Kong, CHINA
       McBride, William J, Boonton, NJ, UNITED STATES
       Gu, Zhengxing, Warren, NJ, UNITED STATES
       Immunomedics, Inc., Morris Plains, NJ, UNITED STATES (U.S. corporation)
PΆ
PΙ
       US 2006140858
                          A1
                               20060629
ΑI
       US 2003-514632
                               20030516 (10)
                          A1
       WO 2003-GB2110
                               20030516
                               20050912 PCT 371 date
       US 2002-10150654
PRAI
                           20020517
DТ
       Utility
FS
       APPLICATION
       HELLER EHRMAN WHITE & MCAULIFFE LLP, 1717 RHODE ISLAND AVE, NW,
LREP
       WASHINGTON, DC, 20036-3001, US
CLMN
       Number of Claims: 127
       Exemplary Claim: 1
ECL
DRWN
       7 Drawing Page(s)
LN.CNT 4528
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention relates to targetable constructs which may be
AB
       bound by a bi-specific antibody or antibody fragment having at least one
       arm that specifically binds a targeted tissue and at least one other arm
       that specifically binds the targetable construct. The targetable
       construct comprises a carrier portion which comprises or bears at least
       one epitope recognizable by at least one arm of said bi-specific
       antibody or antibody fragment. The targetable construct further
       comprises one or more therapeutic or diagnostic agents or enzymes. The
       invention provides constructs and methods for producing the targetable
       constructs and bi-specific antibodies or antibody fragments, as well as
       methods for using them.
    ANSWER 2 OF 22 USPATFULL on STN
L17
AN
       2006:40112 USPATFULL
       Production and use of novel peptide-based agents with bispecific
ΤI
       antibodies
IN
       Goldenberg, David M., Mendham, NJ, UNITED STATES
       Hansen, Hans J., Picayune, MS, UNITED STATES
       Leung, Shui-on, Madison, NJ, UNITED STATES
       McBride, William J., Boonton, NJ, UNITED STATES
       Qu, Zhengxing, Warren, NJ, UNITED STATES
PA
       Immunomedics, Inc., Morris Plains, NJ, UNITED STATES (U.S. corporation)
PΙ
       US 2006034759
                          A1
                               20060216
                               20050808 (11)
ΑI
       US 2005-198846
                          A1
RLI
       Division of Ser. No. US 2002-150654, filed on 17 May 2002, PENDING
       Continuation-in-part of Ser. No. US 1999-382186, filed on 23 Aug 1999,
       PENDING Continuation-in-part of Ser. No. US 2001-823746, filed on 3 Apr
       2001, GRANTED, Pat. No. US 6962702 Continuation-in-part of Ser. No. US
       1999-337756, filed on 22 Jun 1999, PENDING Continuation-in-part of Ser.
      No. US 1999-337756, filed on 22 Jun 1999, PENDING
PRAI
       US 1998-104156P
                           19981014 (60)
       US 1998-90142P
                           19980622 (60)
DT
       Utility
FS
      APPLICATION
LREP
      HELLER EHRMAN WHITE & MCAULIFFE LLP, 1717 RHODE ISLAND AVE, NW,
       WASHINGTON, DC, 20036-3001, US
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CLMN Number of Claims: 95 ECL Exemplary Claim: 1-33

DRWN 7 Drawing Page(s)

LN.CNT 4591

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to a bi-specific antibody or antibody fragment having at least one arm that specifically binds a targeted tissue and at least one other arm that specifically binds a targetable construct. The targetable construct comprises a carrier portion which comprises or bears at least one epitope recognizable by at least one arm of said bi-specific antibody or antibody fragment. The targetable construct further comprises one or more therapeutic or diagnostic agents or enzymes. The invention provides constructs and methods for producing the bi-specific antibodies or antibody fragments, as well as methods for using them.

- L17 ANSWER 3 OF 22 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2006:452992 CAPLUS
- TI An intranasal vaccine targeting both the Bacillus anthracis toxin and bacterium provides protection against aerosol spore challenge in rabbits
- AU Wimer-Mackin, S.; Hinchcliffe, M.; Petrie, C. R.; Warwood, S. J.; Tino, W. T.; Williams, M. S.; Stenz, J. P.; Cheff, A.; Richardson, C.
- CS LigoCyte Pharmaceuticals Inc., Bozeman, MT, 59718, USA
- SO Vaccine (2006), 24(18), 3953-3963 CODEN: VACCDE; ISSN: 0264-410X
- PB Elsevier B.V.
- DT Journal
- LA English
- AB An intranasal vaccine targeting the Bacillus anthracis toxin and vegetative bacterium was tested for the ability to protect immunized rabbits against aerosol B. anthracis spore exposure. Rabbits were vaccinated intranasally with PA-based vaccines formulated as dry powders with or without chitosan (ChiSys®, Archimedes Development Limited), a compound that exhibits muco-adhesive properties, or as a liquid Formulations also contained MPL adjuvant and PA. Some vaccines contained PA conjugated to a 10-mer peptide of the poly--

glutamic acid capsule of B. anthracis.

Rabbits were immunized on days 0 and 28 and aerosol challenged with an average 250 LD50 Ames spores on day 85. Serum antibody was measured before and after challenge. Significant anti-PA serum IgG levels were obtained, particularly with use of ChiSys® based formulations. PA-Conj induced significant anti-capsule responses, although a formulation containing free capsule peptide did not. All immunized rabbits survived the challenge, but differences in morbidity, as evidenced by anorexia, between vaccine groups were observed Only rabbits immunized with PA + PA-Conj appeared normal throughout the post-challenge observation period (14 days), while all that received PA with the free capsule peptide appeared ill at times as evidenced by a failure to eat normally. One neg. control rabbit received a lower inhaled spore dose (183 LD50) and survived the challenge, although it was anorexic post-challenge. It also had a high level of anti-LF antibodies in its convalescent serum (5400 U/mL), indicating an extensive infection. In contrast, 75% of the immunized rabbits had no LF-specific antibody in their post-challenge sera, and the rest had low levels (≤138 U/mL), indicating that infections resulting in toxin production were avoided or greatly reduced. Thus, intranasal immunization with a chitosan-based powder vaccine combining PA and capsule epitopes provided superior protection against B. anthracis infection compared to a single antigen (PA) vaccine, as evidenced by a reduction in morbidity and prevention of death.

RE.CNT 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

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DN
     142:175358
     Anthrax conjugate vaccine induces antibodies to both
ΤI
     bacilli and anthrax toxins
     Wang, Julia Y.; Mekalanos, John; Rhie, Gi-Eun; Collier, John R.
IN
     President and Fellows of Harvard College, USA; The Brigham and Women's
PA
     Hospital, Inc.
SO
     PCT Int. Appl., 29 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
     PATENT NO.
                       KIND
                                DATE
                                          APPLICATION NO.
                                                                  DATE
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     WO 2005007804
                        A2
                                20050127
                                          WO 2004-US10933
                                                                  20040409
PT
     WO 2005007804
                        A3
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            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
            BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
            ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
             SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
             TD, TG
PRAI US 2003-461406P
                         P
                                20030410
     The present invention is directed to immunogenic conjugates
     comprised of poly-\gamma-D-glutamic acid (PGA) covalently bound to
     protective antigen (PA). The invention includes methods for making
     conjugates, vaccines in which they are present, and methods for
     immunizing individuals in which they are used. Conjugation produces a
     synergistic effect dramatically increasing the response of animals to both
     the PGA and PA components of the conjugate. Antibodies to PGA
     and PA confer protection against both bacilli and anthrax
     toxins. The PGA for the vaccine is purified from B. licheniformis and the
     recombinant PA was expressed in E. coli and purified before the synthesis
     of the vaccine conjugate. Also prepared was a PGA-hepatitis B
     core protein conjugate.
    ANSWER 5 OF 22 CAPLUS COPYRIGHT 2006 ACS on STN
L17
AN
     2005:14426 CAPLUS
DN
     142:112426
    Bacillus capsular poly-γ-glutamic acid conjugates for
ΤI
     eliciting immune responses against Bacillus infection
     Schneerson, Rachel; Leppla, Stephen; Robbins, John B.; Shiloach, Joseph;
IN
     Kubler-Kielb, Joanna; Liu, Darrell; Majadly, Fathy
    United States Dept. of Health and Human Services, USA
PA
SO
     PCT Int. Appl., 67 pp.
     CODEN: PIXXD2
DT
     Patent
LA
    English
FAN.CNT 3
    PATENT NO.
                        KIND
                                DATE
                                           APPLICATION NO.
                                                                   DATE
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                                -----
                                            -----
                                                                    -----
PΙ
    WO 2005000884
                         A1
                                20050106
                                            WO 2004-US17736
                                                                    20040604
    WO 2005000884
                         C1
                                20051006
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
            TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
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AN

2005:76327 CAPLUS

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RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
             EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
             SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
             SN, TD, TG
     AU 2004252091
                          A1
                                20050106
                                            AU 2004-252091
                                                                    20040604
                                20050106
                                            CA 2004-2528067
     CA 2528067
                          AA
                                20060315
                                            EP 2004-754360
                                                                    20040604
     EP 1633778
                          A1
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK
                                20051208
                                            US 2004-5851
                                                                    20041206
     US 2005271675
                          A1
                                20051215
                                            WO 2005-US19678
                                                                    20050603
     WO 2005117965
                          A1
             AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
         W:
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
             LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
             NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK,
             SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU,
             ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
             EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT,
             RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
             MR, NE, SN, TD, TG
                                            US 2005-559825
                                20060622
                                                                    20051202
     US 2006134143
                          A1
PRAI US 2003-476598P
                          P
                                20030605
    WO 2004-US17736
                          W
                                20040604
     US 2004-5851
                          Α
                                20041206
     Immunogenic compns. and methods for eliciting an immune response against
AB
     B. anthracis and other bacilli are provided that include
     immunogenic conjugates of a poly-γ-glutamic acid
     (\gamma PGA) polypeptides of B. anthracis, or of another
     Bacillus that expresses a \gammaPGA polypeptide. The \gammaPGA
     conjugates elicit an effective immune response against B.
     anthracis, or against another Bacillus, in mammalian hosts to
     which the conjugates are administered. The conjugate
     consists of γ-D-PGA and carrier selected from bovine serum albumin,
     recombinant Bacillus protective antigen, recombinant Pseudomonas
     aeruginosa exotoxin A, tetanus toxoid, diphtheria toxoid, pertussis
     toxoid, Clostridium perfringens toxoid, HBsAg, HBcAg, heyhole limpet
     hemocyanin, horseshoe crab hemocyanin, edestin, mammalian serum albumin,
     mammalian Ig., analog or mimetic, or combination of two or more. The
     preferred conjugate consists of \gamma-D-PGA and Bacillus
     protective antigen.
RE.CNT 9
              THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L17
     ANSWER 6 OF 22 USPATFULL on STN
AN
       2005:275142 USPATFULL
TI
       Compositions and methods for topical diagnostic and therapeutic
       transport
       Dake, Michael D., Stanford, CA, UNITED STATES
IN
       Waugh, Jacob M., Mountain View, CA, UNITED STATES
       Essentia Biosystems, Inc, Mountain View, CA, UNITED STATES (U.S.
PA
       corporation)
PΙ
       US 2005239705
                          A1
                               20051027
ΑI
       US 2005-73307
                          A1
                               20050303 (11)
PRAI
       US 2004-550014P
                           20040303 (60)
DT
       Utility
FS
       APPLICATION
LREP
      MORGAN & FINNEGAN, L.L.P., 3 World Financial Center, New York, NY,
       10381-2101, US
CLMN
      Number of Claims: 191
ECL
      Exemplary Claim: 1
```

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CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Compositions and methods are provided that are useful for the delivery,
       including transdermal delivery, of biologically active agents, such as
       non-protein non-nucleotide therapeutics and protein-based therapeutics
       excluding insulin, botulinum toxins, antibody fragments, and VEGF. The
       compositions and methods are particularly useful for topical delivery of
       antifungal agents and antigenic agents suitable for immunization.
       Alternately, the compositions can be prepared with components useful for
       targeting the delivery of the compositions as well as imaging
       components.
L17 ANSWER 7 OF 22 USPATFULL on STN
AN
       2005:157794 USPATFULL
ΤI
       Fluorinated carbohydrate conjugates
IN
       McBride, William J., Boonton, NJ, UNITED STATES
       Goldenberg, David M., Mendham, NJ, UNITED STATES
PA
       Immunomedics, Inc., Morris Plains, NJ, UNITED STATES (U.S. corporation)
PΙ
       US 2005136001
                          A1
                               20050623
       US 2004-901441
AΙ
                          A1
                               20040729 (10)
       US 2003-490884P
PRAI
                           20030729 (60)
DT
       Utility
       APPLICATION
FS
LREP
       Paul M. Booth, Ph.D., Heller Ehrman White & McAuliffe, 1717 Rhode Island
       Avenue, N.W., Washington, DC, 20036-3001, US
CLMN
       Number of Claims: 18
ECL
       Exemplary Claim: 1
DRWN
       15 Drawing Page(s)
LN.CNT 2380
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Disclosed are novel conjugates that include fluorinated
       carbohydrate molecules and methods for synthesizing the
       conjugates. The fluorinated carbohydrate molecule may include a
       radioisotope. The method of synthesizing the conjugate is
       useful for labeling selected molecules, and the conjugates may
       be useful in diagnostic or therapeutic methods. Particularly, the
       conjugates may be useful in diagnostic or therapeutic kits.
    ANSWER 8 OF 22 USPATFULL on STN
L17
AN
       2005:138029 USPATFULL
TI
       Modified polypeptides with therapeutic activity and methods of use
       Mayo, Kevin H., Minneapolis, MN, UNITED STATES
TN
       Regents of the University of Minnesota, Minneapolis, MN, UNITED STATES
PΑ
       (U.S. corporation)
PT
       US 2005118678
                               20050602
                          A1
       US 2004-967060
AΤ
                          Α1
                               20041015 (10)
       US 2003-512372P
PRAI
                           20031017 (60)
DТ
       Utility
FS
       APPLICATION
       MUETING, RAASCH & GEBHARDT, P.A., P.O. BOX 581415, MINNEAPOLIS, MN,
LREP
       55458, US
CLMN
       Number of Claims: 44
ECL
       Exemplary Claim: 1
DRWN
       16 Drawing Page(s)
LN.CNT 2102
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Polypeptides modified by fatty acid conjugation and methods of using
       such modified polypeptides in treating bacterial infections, including
       the treatment of antibiotic resistant bacterial infections, are
       disclosed.
     ANSWER 9 OF 22 USPATFULL on STN
L17
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DRWN

LN.CNT 3191

17 Drawing Page(s)

2005:117278 USPATFULL

AN

```
Multivalent carriers of bi-specific antibodies
TT
       Hansen, Hans J., Picayune, MS, UNITED STATES
TN
       McBride, William J., Boonton, NJ, UNITED STATES
       Qu, Zhengxing, Warren, NJ, UNITED STATES
       Immunomedics, Inc., Morris Plains, NJ, UNITED STATES (U.S. corporation)
PA
PΙ
       US 2005100543
                          A1
                               20050512
       US 2004-882151
                          A1
                               20040701 (10)
ΑI
      US 2003-483832P
                           20030701 (60)
PRAI
DT
      Utility
       APPLICATION
FS
       HELLER EHRMAN WHITE & MCAULIFFE LLP, 1717 RHODE ISLAND AVE, NW,
LREP
       WASHINGTON, DC, 20036-3001, US
       Number of Claims: 35
CLMN
ECL
       Exemplary Claim: 1
DRWN
       9 Drawing Page(s)
LN.CNT 5871
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Provided herein are targetable constructs that are multivalent carriers
AΒ
       of bi-specific antibodies, i.e., each molecule of a targetable construct
       can serve as a carrier of two or more bi-specific antibodies. Also
       provided are targetable complexes formed by the association of a
       targetable construct with two or more bi-specific antibodies. The
       targetable constructs and targetable complexes of the invention are
       incorporated into biosensors, kits and pharmaceutical compositions, and
       are used in a variety of therapeutic and other methods.
     ANSWER 10 OF 22 USPATFULL on STN
L17
       2005:30279 USPATFULL
AN
ΤI
       D-amino acid peptides
       McBride, William J., Boonton, NJ, UNITED STATES
IN
       Goldenberg, David M., Mendham, NJ, UNITED STATES
       Immunomedics, Inc., Morris Plains, NJ (U.S. corporation)
PA
ΡI
       US 2005025709
                          A1
                               20050203
       US 2004-866180
                               20040614 (10)
ΑI
                          Α1
PRAI
       US 2003-478403P
                           20030613 (60)
       Utility
DT
FS
       APPLICATION
       HELLER EHRMAN WHITE & MCAULIFFE LLP, 1666 K STREET, NW, SUITE 300,
LREP
       WASHINGTON, DC, 20006
CLMN
       Number of Claims: 151
ECL
       Exemplary Claim: 1
DRWN
       18 Drawing Page(s)
LN.CNT 4255
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB
       The present invention provides compounds of the formula
       X--R.sup.1-D-[Dpr, Orn or Lys] (A) -R.sup.2(Z) -D-[Dpr, Orn or
       Lys] (B) -- R. sup. 3 (Y) -- NR. sup. 4R. sup. 5; or R. sup. 1 (X) - D- [Dpr, Orn or
       Lys] (A) -R.sup.2(Z) -D-[Dpr, Orn or Lys](B) --R.sup.3(Y) --NR.sup.4R.sup.5,
       in which X is a hard acid cation chelator, a soft acid cation chelator
       or Ac--, R.sup.1, R.sup.2 and R.sup.3 are independently selected from a
       covalent bond or one or more D-amino acids that can be the same or
       different, Y is a hard acid cation chelator, a soft acid cation chelator
       or absent, Z is a hard acid cation chelator, a soft acid cation chelator
       or absent, and A and B are haptens or hard acid cation chelators and can
       be the same or different, and R.sup.4 and R.sup.5 are independently
       selected from the group consisting of hard acid cation chelators, soft
       acid cation chelators, enzymes, therapeutic agents, diagnostic agents
       and H. The present invention also provides methods of using these
       compounds and kits containing the compounds.
L17 ANSWER 11 OF 22 USPATFULL on STN
```

AN

TI

2005:3843 USPATFULL

multispecific antibodies

Therapeutic and diagnostic conjugates for use with

```
IN
       McBride, William J., Boonton, NJ, UNITED STATES
       Goldenberg, David M., Mendham, NJ, UNITED STATES
       Noren, Carl, Mt. Arlington, NJ, UNITED STATES
       Hansen, Hans J., Picayune, MS, UNITED STATES
       IMMUNOMEDICS, INC. (U.S. corporation)
PA
PΙ
       US 2005002945
                          A1
                               20050106
AΙ
       US 2004-776470
                          A1
                               20040211 (10)
       Continuation-in-part of Ser. No. US 2002-150654, filed on 17 May 2002,
RLI
       PENDING Continuation-in-part of Ser. No. US 1999-382186, filed on 23 Aug
       1999, ABANDONED Continuation-in-part of Ser. No. US 1999-337756, filed
       on 22 Jun 1999, PENDING Continuation-in-part of Ser. No. US 2001-823746,
       filed on 3 Apr 2001, PENDING Continuation-in-part of Ser. No. US
       1999-337756, filed on 22 Jun 1999, PENDING
PRAI
      US 1998-90142P
                          19980622 (60)
      US 1998-104156P
                           19981014 (60)
      US 1998-90142P
                           19980622 (60)
       US 1998-104156P
                           19981014 (60)
DT
       Utility
       APPLICATION
FS
LREP
       FOLEY AND LARDNER, SUITE 500, 3000 K STREET NW, WASHINGTON, DC, 20007
CLMN
      Number of Claims: 135
ECL
       Exemplary Claim: 1
DRWN
       18 Drawing Page(s)
LN.CNT 3522
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Disclosed are compounds that include two or more haptens
       conjugated by a spacer or a carrier. The haptens may include
       diethylenetriaminepentaacetate (DTPA), histimine-succinyl-glutamine
       (HSG), or combinations of DTPA and HSG. The compound also includes an
       effector molecule which may be conjugated to one or more of
       the haptens, the spacer/carrier, or both. The effector molecule may be
       conjugated by a number of linkages including an ester linkage,
       an imino linkage, an amino linkage, a sulfide linkage, a
       thiosemicarbazone linkage, a semicarbazone linkage, an oxime linkage, an
       ether linkage, or combinations of these linkages. Also disclosed are
       methods of synthesizing the compounds and/or precursors of the
       compounds.
    ANSWER 12 OF 22 USPATFULL on STN
L17
       2004:280799 USPATFULL
AN
       Multi-component biological transport systems
TI
       Waugh, Jacob, Mountain View, CA, UNITED STATES
IN
       Dake, Michael, Stanford, CA, UNITED STATES
       Essentia Biosystems, Inc., Mountain View, CA (U.S. corporation)
PA
PΙ
       US 2004220100
                          A1
                               20041104
                               20040303 (10)
ΑI
       US 2004-793138
                          A1
       Continuation-in-part of Ser. No. US 2001-910432, filed on 20 Jul 2001,
RLI
       PENDING
PRAI
       US 2000-220244P
                           20000721 (60)
DT
      Utility
FS
       APPLICATION
LREP
       MORGAN & FINNEGAN, L.L.P., 3 WORLD FINANCIAL CENTER, NEW YORK, NY,
       10281-2101
CLMN
       Number of Claims: 240
ECL
       Exemplary Claim: 1
DRWN
       12 Drawing Page(s)
LN.CNT 3742
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Compositions and methods are provided that are useful for the delivery,
       including transdermal delivery, of biologically active agents, including
       nucleic acids and therapeutic proteins including insulin, larger
       therapeutic proteins such as botulinum toxin and other biologically
       active agents such as a therapeutic protein which does not
       therapeutically alter blood glucose levels, a therapeutic nucleic
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acid-based agent, a non-protein non-nucleic acid therapeutic agent such as an antifungal agent or alternately an agent for immunization. The compositions can be prepared with components useful for targeting the delivery of the compositions as well as imaging components.

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L17 ANSWER 13 OF 22 USPATFULL on STN
AN
       2004:203967 USPATFULL
ΤI
       Pyrrolidones with anti-HIV activity
IN
       Wu, Baogen, San Diego, CA, UNITED STATES
       He, Yun, San Diego, CA, UNITED STATES
       Ngyuen, Truc, San Diego, CA, UNITED STATES
       Kuhen, Kelli L., Carlsbad, CA, UNITED STATES
       Ellis, David Archer, San Diego, CA, UNITED STATES
       Jiang, Tao, San Diego, CA, UNITED STATES
       Xe, Xiaohui, San Diego, CA, UNITED STATES
       Yang, Kunyong, San Diego, CA, UNITED STATES
       Bursulaya, Badry, San Diego, CA, UNITED STATES
PA
       IRM LLC, a Delaware LLC, Hamilton HM LX, BERMUDA (U.S. corporation)
PΙ
       US 2004157859
                         A1
                               20040812
AΙ
      US 2003-690873
                          A1
                               20031021 (10)
PRAI
      US 2002-422619P
                           20021030 (60)
      US 2002-420480P
                           20021021 (60)
DT
      Utility
FS
       APPLICATION
       TOWNSEND AND TOWNSEND AND CREW, LLP, TWO EMBARCADERO CENTER, EIGHTH
LREP
       FLOOR, SAN FRANCISCO, CA, 94111-3834
CLMN
      Number of Claims: 22
ECL
       Exemplary Claim: 1
       101 Drawing Page(s)
DRWN
LN.CNT 3331
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention relates to inhibition of viruses, e.g., HIV using
       pyrrolidones and compounds related to pyrrolidones. The invention
       further relates to methods for identifying and using agents, including
       small molecule chemical compositions that inhibit HIV in a cell; as well
       as to methods of prophylaxis, and therapy related to HIV infection and
       related disease states such as AIDS.
L17 ANSWER 14 OF 22 USPATFULL on STN
       2004:197449 USPATFULL
AN
       Oxindoles with anti-HIV activity
ΤI
      He, Yun, San Diego, CA, UNITED STATES
IN
       Jiang, Tao, San Diego, CA, UNITED STATES
       Kuhen, Kelli L., Carlsbad, CA, UNITED STATES
       Ellis, David Archer, San Diego, CA, UNITED STATES
      Wu, Baogen, San Diego, CA, UNITED STATES
      Wu, Tom Yao-Hsiang, La Jolla, CA, UNITED STATES
       Bursulaya, Badry, San Diego, CA, UNITED STATES
PA
       IRM LLC, a Delaware LLC, Hamilton, BERMUDA (U.S. corporation)
PΙ
       US 2004152755
                          A1
                               20040805
ΑI
      US 2003-690802
                          A1
                               20031021 (10)
PRAI
       US 2002-420482P
                           20021021 (60)
       US 2002-420481P
                           20021021 (60)
DT
       Utility
       APPLICATION
FS
LREP
       TOWNSEND AND TOWNSEND AND CREW, LLP, TWO EMBARCADERO CENTER, EIGHTH
       FLOOR, SAN FRANCISCO, CA, 94111-3834
      Number of Claims: 17
CLMN
ECL
       Exemplary Claim: 1
       21 Drawing Page(s)
DRWN
LN.CNT 2180
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB
       The present invention relates to inhibition of viruses, e.g., HIV using
       oxindoles and compounds related to oxindoles. The invention further
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relates to methods for identifying and using agents, including small molecule chemical compositions that inhibit HIV in a cell; as well as to methods of prophylaxis, and therapy related to HIV infection and related disease states such as AIDS.

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L17 ANSWER 15 OF 22 USPATFULL on STN
       2004:197413 USPATFULL
AN
ΤI
       Quinolones with anti-HIV activity
IN
       He, Yun, San Diego, CA, UNITED STATES
       Ellis, David Archer, San Diego, CA, UNITED STATES
       Anaclerio, Beth Marie, San Diego, CA, UNITED STATES
       Kuhen, Kelli L., Carlsbad, CA, UNITED STATES
       Wu, Baogen, San Diego, CA, UNITED STATES
       Jiang, Tao, San Diego, CA, UNITED STATES
       IRM LLC, a Delaware LLC, Hamilton, HM LX, BERMUDA (U.S. corporation)
PΑ
                               20040805
PΙ
       US 2004152719
                         A1
       US 7019141
                          B2
                               20060328
ΑI
       US 2003-690738
                          A1
                               20031021 (10)
PRAI
       US 2002-420163P
                          20021021 (60)
DT
       Utility
FS
       APPLICATION
       TOWNSEND AND TOWNSEND AND CREW, LLP, TWO EMBARCADERO CENTER, EIGHTH
LREP
       FLOOR, SAN FRANCISCO, CA, 94111-3834
       Number of Claims: 17
CLMN
ECL
       Exemplary Claim: 1
DRWN
       14 Drawing Page(s)
LN.CNT 2027
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention relates to inhibition of viruses, e.g., HIV using
       quinolones and compounds related to quinolones. The invention further
       relates to methods for identifying and using agents, including small
       molecule chemical compositions that inhibit HIV in a cell; as well as to
       methods of prophylaxis, and therapy related to HIV infection and related
       disease states such as AIDS.
L17 ANSWER 16 OF 22 USPATFULL on STN
AN
       2004:76181 USPATFULL
ΤI
       Immunogenicity-enhancing carriers and compositions thereof and methods
       of using the same
TN
       Waggoner, David W., JR., Seattle, WA, UNITED STATES
       Coon, Michael E., Seattle, WA, UNITED STATES
PΙ
       US 2004057958
                         A1
                               20040325
ΑI
       US 2003-441944
                               20030519 (10)
                          A1
       US 2002-381550P
PRAI
                           20020517 (60)
DT
       Utility
FS
       APPLICATION
LREP
       DONALD W. WYATT, CELL THERAPEUTICS, INC., 501 ELLIOTT AVENUE WEST, #400,
       SEATTLE, WA, 98119
CLMN
       Number of Claims: 85
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 2556
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention is directed to compositions comprising a
       substantially non-antigenic carrier associated with an antigen and the
       use of such compositions to enhance the immunogenicity of the associated
       antigen. In addition, the compositions of the invention may be used to
       generate an immune response directed predominantly to an antigen
       associated with a carrier. Specific carriers of the invention include
       homopolymers and copolymers of polyamino acids. Compositions of the
       invention are used according to the invention to elicit or enhance an
       immune response directed against an antigen and may be used for the
       prevention and treatment of infection and disease, for example.
      Additionally, compositions of the invention are useful for generating an
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antibodies specific for an antigen and, accordingly, may be used to generate antigen-specific antibodies suitable for the diagnosis or treatment of infection and disease.

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1.17
    ANSWER 17 OF 22 USPATFULL on STN
AN
       2004:70645 USPATFULL
       Electroporation methods for introducing bioactive agents into cells
ΤI
       Barman, Shikha P., Bedford, MA, UNITED STATES
IN
       Hedley, Mary Lynne, Lexington, MA, UNITED STATES
       Wang, Daqing, Bedford, MA, UNITED STATES
                               20040318
PΙ
       US 2004053873
                          A1
       US 2003-370131
                          A1
                               20030219 (10)
AΙ
                           20020215 (60)
PRAI
      US 2002-357542P
      Utility
DT
FS
      APPLICATION
      FISH & RICHARDSON PC, 225 FRANKLIN ST, BOSTON, MA, 02110
LREP
CLMN
      Number of Claims: 33
       Exemplary Claim: 1
ECL
DRWN
       6 Drawing Page(s)
LN.CNT 1987
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention provides compositions and methods for introducing
AΒ
       bioactive agents into cells. Bioactive agents are provided together with
       a delivery vehicle and a cell is subjected to electroporation, thereby
       resulting in the introduction of the bioactive agent into the cell.
    ANSWER 18 OF 22 USPATFULL on STN
L17
       2004:57028 USPATFULL
AN
       Polymeric delivery systems
TТ
       Griffiths, Gary L., Morristown, NJ, UNITED STATES
IN
       Goldenberg, David M., Medham, NJ, UNITED STATES
       Hansen, Hans J., Picayune, MS, UNITED STATES
       Immunomedics, Inc. (U.S. corporation)
PA
       US 2004043030
PΤ
                          A1
                               20040304
                               20030609 (10)
ΑI
       US 2003-456580
                          A1
       Continuation-in-part of Ser. No. US 2002-209592, filed on 31 Jul 2002,
RLI
       PENDING
       US 2001-308605P
                           20010731 (60)
PRAI
DT
       Utility
FS
       APPLICATION
       Stephen B. Maebius, Foley & Lardner, Washington Harbour, 3000 K Street,
LREP
       N.W., Suite 500, Washington, DC, 20007-5143
CLMN
       Number of Claims: 73
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 2547
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention relates to a method of targeting an agent towards
       a targeting site in a tissue comprising administering a multi-specific
       antibody or antibody fragment comprising a targeting arm and a capture
       arm that binds to a polymer conjugate, and administering a
       polymer conjugate to the tissue. The present invention also
       relates to a kit for targeting a target site within a comprising a
       multi-specific antibody or antibody fragment comprising a targeting arm
       and a capture arm that binds to a polymer conjugate, and a
       polymer conjugate.
L17 ANSWER 19 OF 22 USPATFULL on STN
       2003:282254 USPATFULL
AN
TI
       Use of bi-specific antibodies for pre-targeting diagnosis and therapy
IN
       Goldenberg, David M., Mendham, NJ, UNITED STATES
       Hansen, Hans J., Picayune, MS, UNITED STATES
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Leung, Shui-On, Shatin, NT, HONG KONG

McBride, William J., Boonton, NJ, UNITED STATES

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Qu, Zhengxing, Warren, NJ, UNITED STATES
PA
       IMMUNOMEDICS, INC. (U.S. corporation)
PΙ
       US 2003198595
                          A1
                               20031023
ΑI
       US 2002-150654
                          A1
                               20020517 (10)
RLI
       Continuation-in-part of Ser. No. US 1999-382186, filed on 23 Aug 1999,
       PENDING Continuation-in-part of Ser. No. US 2001-823746, filed on 3 Apr
       2001, PENDING
PRAI
       US 1998-104156P
                           19981014 (60)
       US 1998-90142P
                           19980622 (60)
       Utility
DT
       APPLICATION
FS
       FOLEY AND LARDNER, SUITE 500, 3000 K STREET NW, WASHINGTON, DC, 20007
LREP
CLMN
       Number of Claims: 127
ECL
       Exemplary Claim: 1
DRWN
       7 Drawing Page(s)
LN.CNT 4670
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention relates to a bi-specific antibody or antibody
       fragment having at least one arm that specifically binds a targeted
       tissue and at least one other arm that specifically binds a targetable
       construct. The targetable construct comprises a carrier portion which
       comprises or bears at least one epitope recognizable by at least one arm
       of said bi-specific antibody or antibody fragment. The targetable
       construct further comprises one or more therapeutic or diagnostic agents
       or enzymes. The invention provides constructs and methods for producing
       the bi-specific antibodies or antibody fragments, as well as methods for
       using them.
    ANSWER 20 OF 22 USPATFULL on STN
L17
       2003:158943 USPATFULL
AN
ΤI
       Therapeutic uses of polyvalent compositions in infectious diseases
IN
       Mekalanos, John J., Charlestown, MA, UNITED STATES
       Wang, Ying, Brookline, MA, UNITED STATES
       Collier, R. John, Wellesley Hills, CA, UNITED STATES
       Mourez, Michael, Boston, MA, UNITED STATES
PΤ
       US 2003108556
                               20030612
                          A1
       US 2002-165762
                               20020607 (10)
AΙ
                          A1
       US 2001-296942P
                           20010608 (60)
PRAI
DT
       Utility
FS
       APPLICATION
       EDWARDS & ANGELL, LLP, P.O. BOX 9169, BOSTON, MA, 02209
LREP
CLMN
       Number of Claims: 48
ECL
       Exemplary Claim: 1
DRWN
       4 Drawing Page(s)
LN.CNT 1705
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB
       New therapeutic methods and compositions are provided for treating
       against an infectious agent in a mammal by administration of a polymeric
       material having linked thereto a plurality of therapeutic agents against
       the infective agent, wherein the polymer comprises polymerized dextran
       or ethylene glycol units. The compositions and methods of the invention
       are particularly useful to treat against bacterial infections, including
       treatment of mammalian cells infected with gram-negative bacteria or
       gram-positive bacteria. The compositions of the invention can be useful
       for treating against anthrax, staphylococcus, pneumococcus and
       other bacteria, parasites, fungi, viral and protozoan infections.
    ANSWER 21 OF 22 USPATFULL on STN
AN
       2002:219333 USPATFULL
TI
       Use of high density microparticles for removal of pathogens
IN
       Cook, David N., Lafayette, CA, UNITED STATES
      Monroy, Rodney L., Rockport, MA, UNITED STATES
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PΙ

US 2002117453

US 6730230

A1

B2

20020829

20040504

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US 2002-43471
                        A1
                               20020111 (10)
AΙ
       US 2001-262443P
PRAI
                         20010116 (60)
DT
       Utility
FS
       APPLICATION
       CARELLA, BAYRNE, BAIN, GILFILLAN,, CECCHI, STEWART & OLSTEIN, 6 Becker
LREP
       Farm Road, Roseland, NJ, 07068
CLMN
       Number of Claims: 32
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 877
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB
       Methods of using high-density microparticles to bind and remove
       pathogens from biological fluids are disclosed. Pathogens include
       prions, viruses, bacteria and protozoa.
L17 ANSWER 22 OF 22 USPATFULL on STN
       1999:50841 USPATFULL
AN
TI
       Antigen-processing cell-targeted conjugates
       Swadesh, Joel K., 285 Plantation St., No. 718, Worcester, MA, United
IN
       States 01604
       Sevoian, Martin, 167 Montague Rd., North Amherst, MA, United States
       01059
PΙ
       US 5898033
                               19990427
ΑI
       US 1997-994334
                               19971219 (8)
RLI
       Continuation of Ser. No. US 1995-475528, filed on 7 Jun 1995, now
       abandoned
DT
       Utility
       Granted
FS
EXNAM Primary Examiner: MacMillan, Keith D.
       Fish & Richardson P.C.
LREP
CLMN
       Number of Claims: 18
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 561
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       An anti-inflammatory conjugate including a polyamino acid
AB
       backbone, a non-steroidal anti-inflammatory agent, and a moiety linking
       the anti-inflammatory agent to the backbone, wherein the polyamino acid
       backbone has a molecular weight greater than 250 kD.
=> s 117 and ((poly glutamic acid?)(2w)(anthrax or anthracis))
             0 L17 AND ((POLY GLUTAMIC ACID?)(2W)(ANTHRAX OR ANTHRACIS))
=> s l17 and ((poly glutamic acid?)(2w)(conjugate?))
             1 L17 AND ((POLY GLUTAMIC ACID?)(2W)(CONJUGATE?))
=> d
L19
     ANSWER 1 OF 1 CAPLUS COPYRIGHT 2006 ACS on STN
AN
     2005:14426
                CAPLUS
DN
     142:112426
ΤI
     Bacillus capsular poly-γ-glutamic acid conjugates for
     eliciting immune responses against Bacillus infection
IN
     Schneerson, Rachel; Leppla, Stephen; Robbins, John B.; Shiloach, Joseph;
     Kubler-Kielb, Joanna; Liu, Darrell; Majadly, Fathy
PA
     United States Dept. of Health and Human Services, USA
SO
     PCT Int. Appl., 67 pp.
     CODEN: PIXXD2
DT
    Patent
LA
    English
FAN.CNT 3
    PATENT NO.
                       KIND
                                DATE
                                           APPLICATION NO.
                                                                   DATE
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             MR, NE, SN, TD, TG
                                20060622
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                          A1
PRAI US 2003-476598P
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                                20030605
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RE.CNT 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT